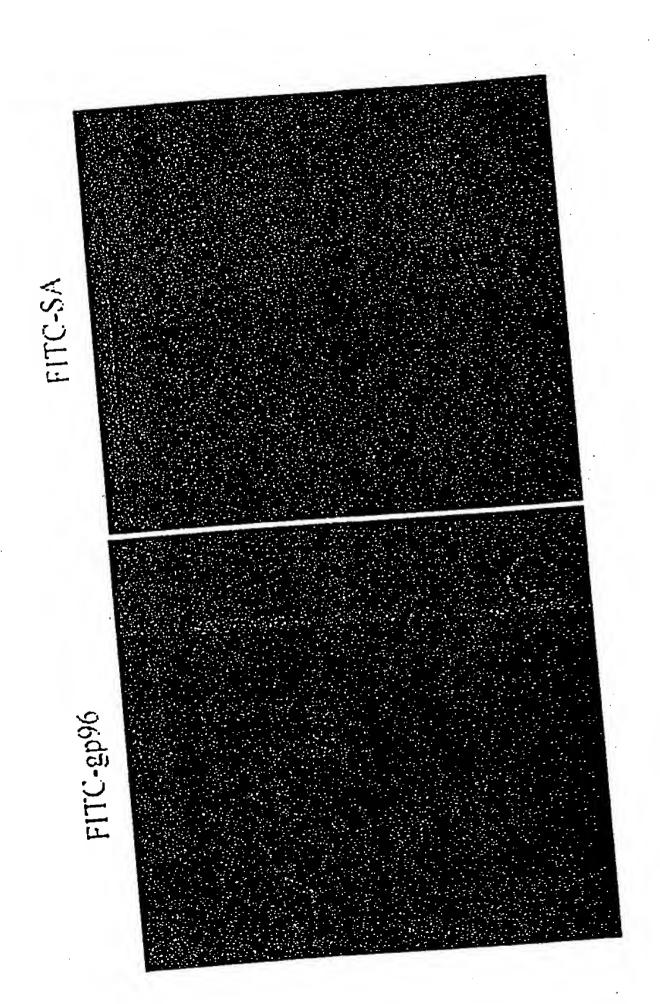


- 12 - White the desired and the second of



8449-134



=1G. 1a

Membranes from	RAW	264.7	P815	
Affinity column	gp96	SA	gp96	<u>.</u>
212 =	1\$		· . •	
116 ⊭				15,100
83 ⊭			÷.	للمرة متولاياك
51 ⊭				•
35 ⊭				
- 28 ⊭				

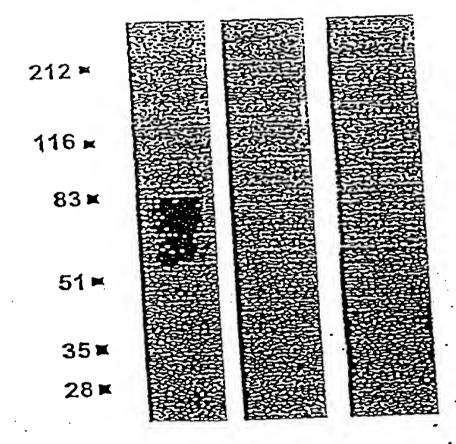
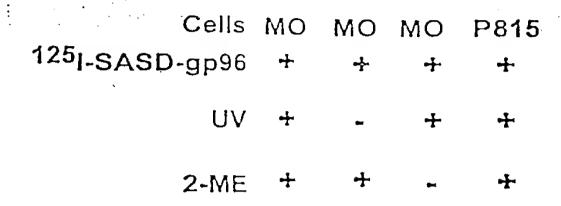


FIG. 1b



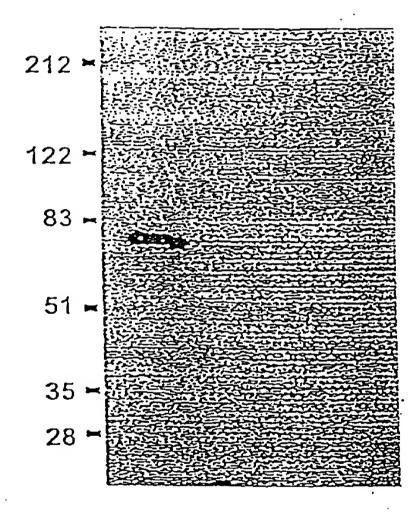


FIG. 1c

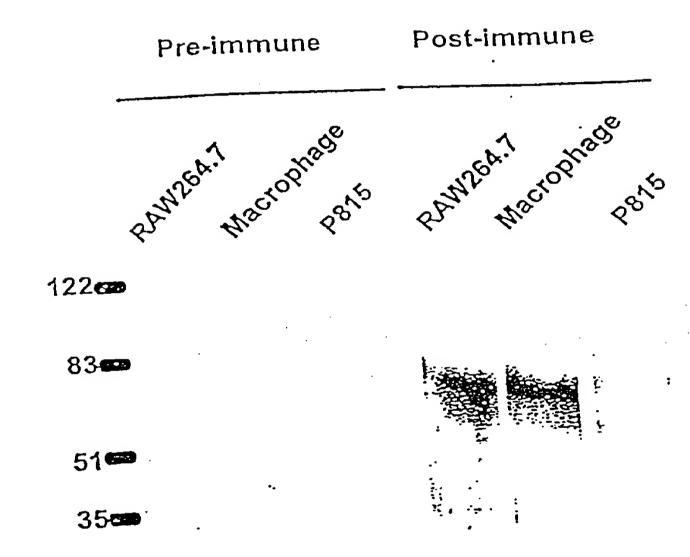


FIG. 2a

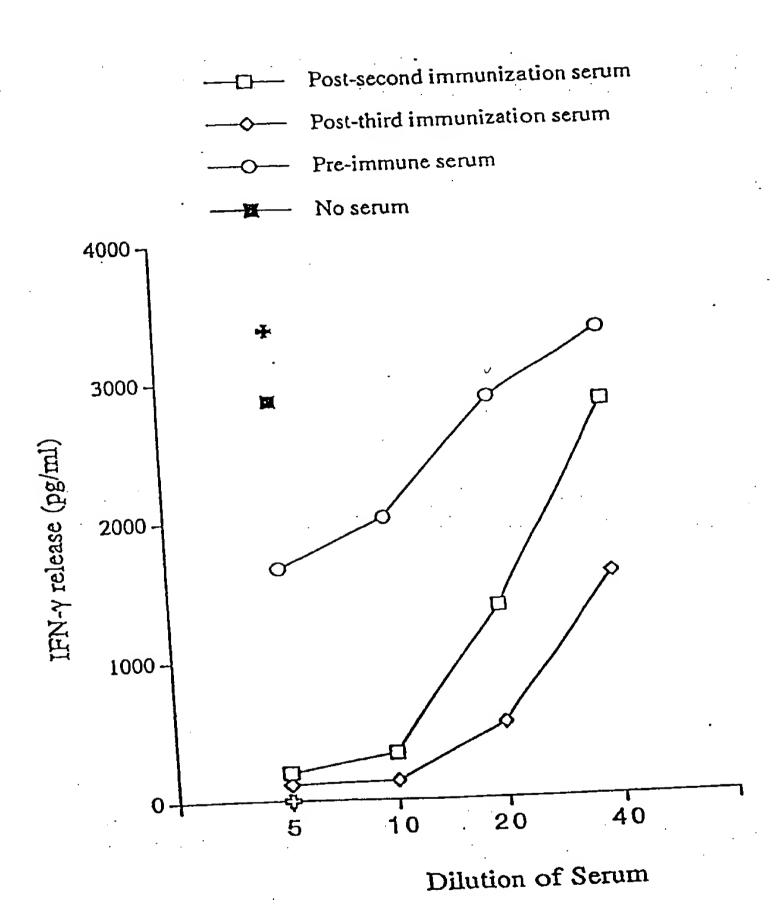


FIG. 2b

Sec] #	b	У	+1
G	1	58.1	-	10
G	2	115.1	1095.2	9
Α	3	186.2	1038.2	8 ·
L	4	299.3	967.1	7
H	5	436.5	853.9	[~] 6
I	6	549.6	716.8	5
Y	7	712.8	603.6	4
H	8	850.0	440.5	3
Q	9	978.1	303.3	2
R	10	- .	175.2	1

FIG. 3a

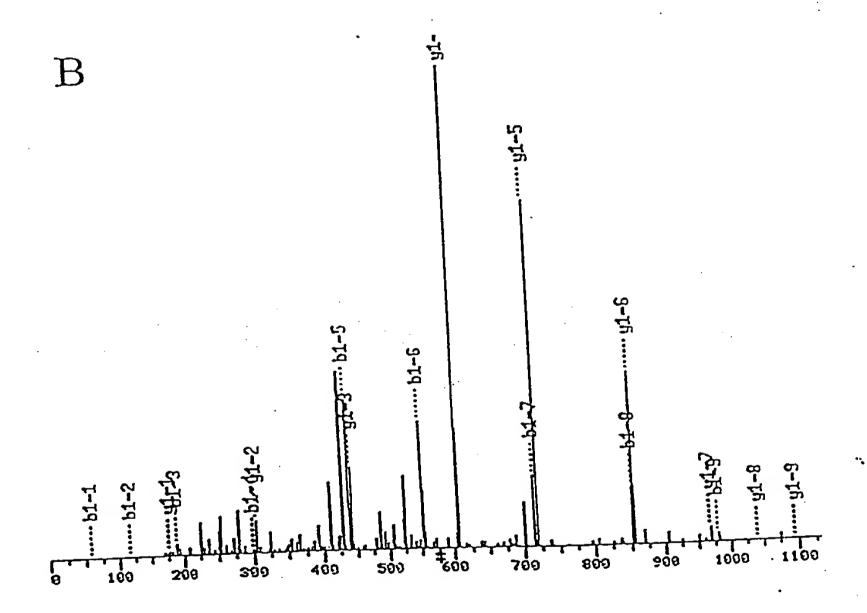
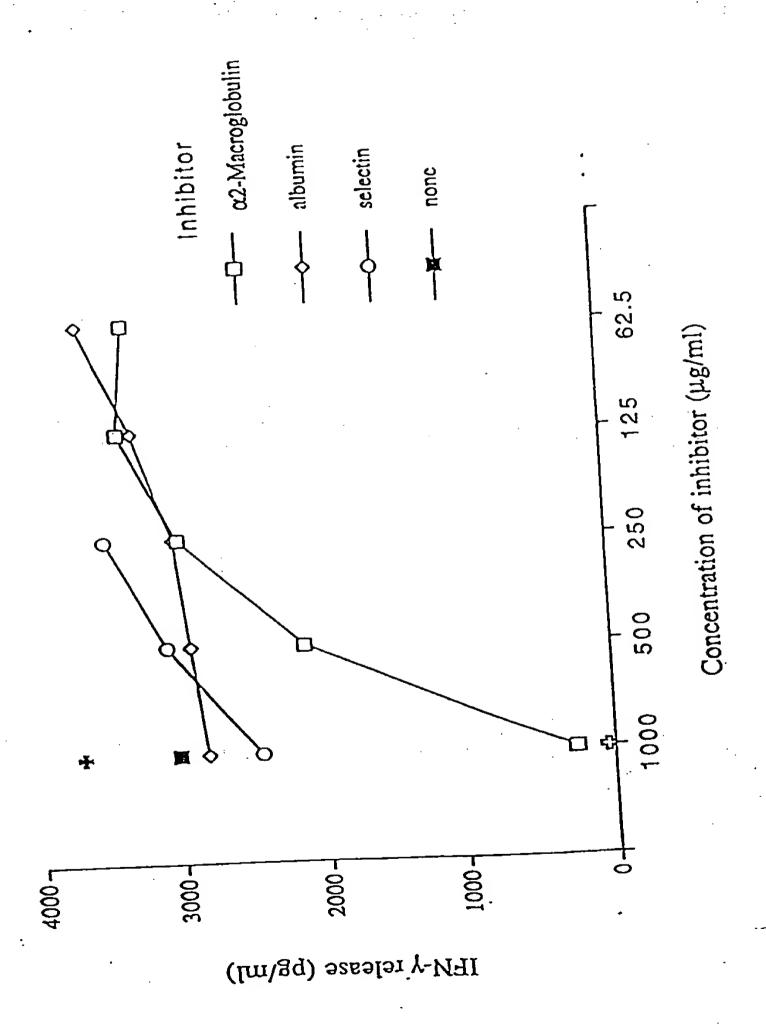


FIG. 3b

Position	•	Sequence
509-518 328-337 460-469 338-348	973.1753	SGFSLGSDGK (Sea 10 M:54) GIALDPAMGK (Sea 10 Mo:54) GGALHIYHQR (Sea 10 Mo:56) VFFTDYGQIPK (Sea 10 Mo:57)
		



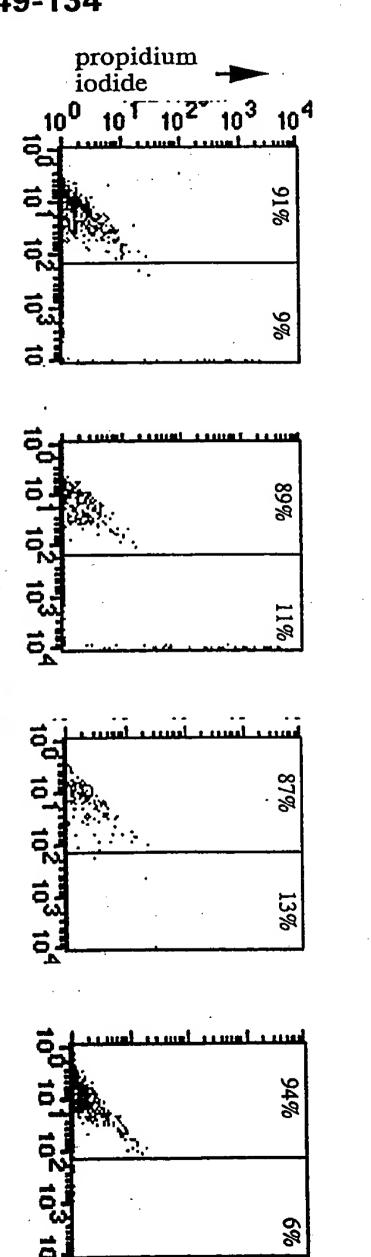
下16. 4

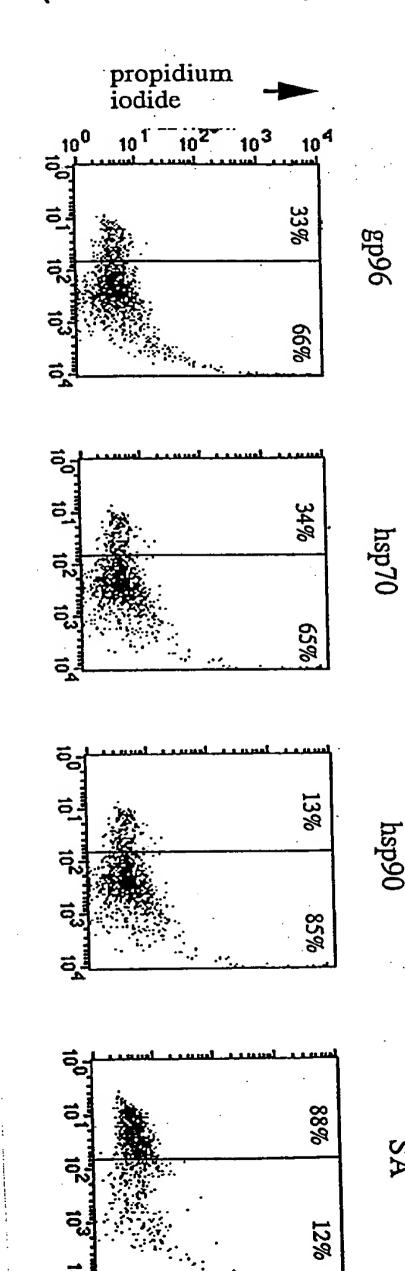
Table 1. Specific binding of HSPs and α_2 -macroglobulin to primary cultures and cell lines of several histological origins*

			**% cells binding with FITC-labeled:								
Cells	Cell type	Haplotype	α₂M	gp96	hsp70	hsp90	SA				
B16	Melanoma	· b	0.1	3.5	6.4	8.0	0.3				
CT26	Carcinoma	d	N/D	0.3	3.1	5.5	0.4				
YAC-1	Lymphoma	b	0.1	3.1	23.0	5.0	0.2				
EL4	T cell thymoma	b	0.1	2.9	3.0	6.6	1.0				
Meth A	Sarcoma	d	0.1	0.1	1.5	0.9	0.5				
PS-C3H	Fibrosarcoma	k	0.1	0.1	2.0	0.3	0.3				
UV6139	Sarcoma	k	11	0.0	0.7	0.2	1.5				
P815	Mastocytoma	d	0.1	1.1	1.7	0.7	0.2				
Peritoneal cells	Macrophage	d	90	97	82	82	11				
BM-DCs	Dendritic cells	b and d	+++#	+++	+++	+++	_				
RAW264.7*	Macrophage	d	76	82	85	90	8.0				
RAW309Cr.1*	Macrophage	bxd	0.1	0.1	0.1	0.1	0.1				

6B

(Sheet 11 of 91)





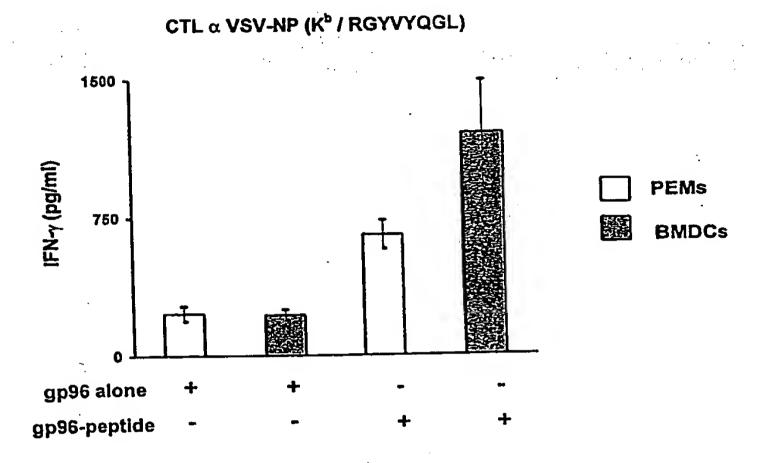


FIG. 7A

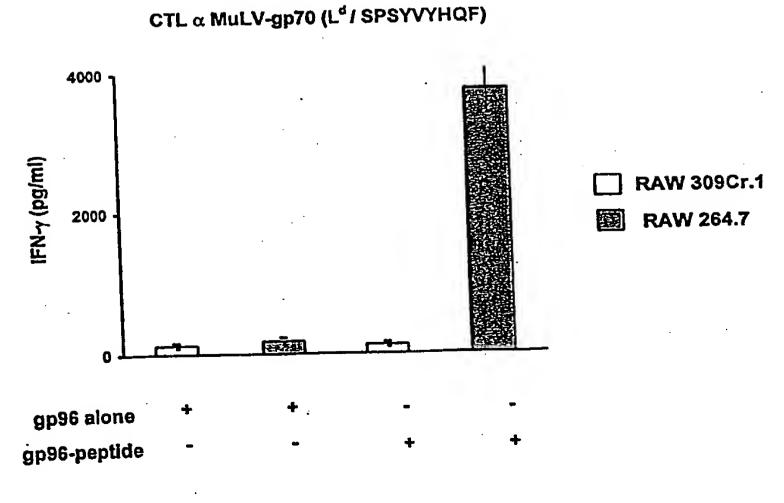


FIG. 7B

gp96 CTL & MuLV-gp70 (L⁴/ SPSYVYHQF) APC: RAW 264.7 hsp90 hsp70 CRT 7500 ₁ - 2000 2500 -Chaperone-AH1/19 Chaperone AH1/19

APC: RAW 264.7 CTL against AH1 (Ld / SPSYVYHQF)

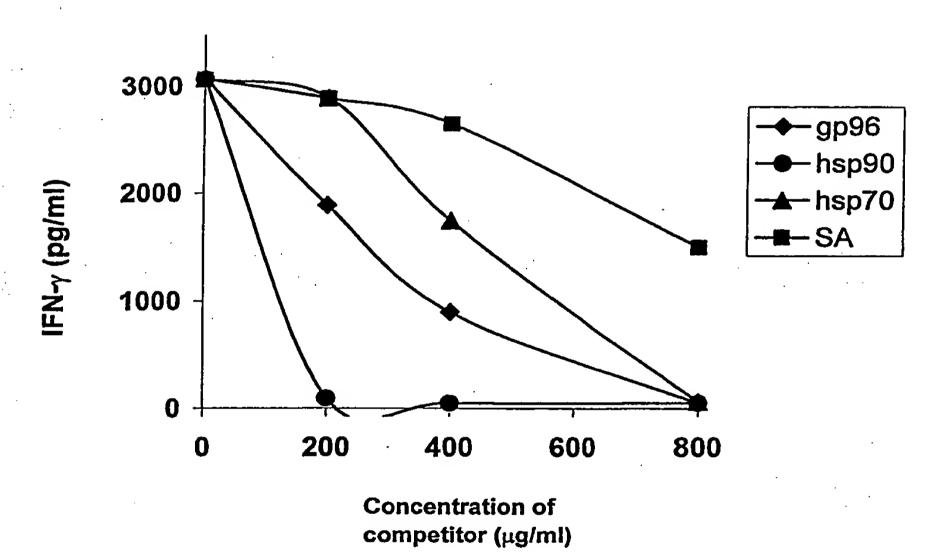


FIG. 9A



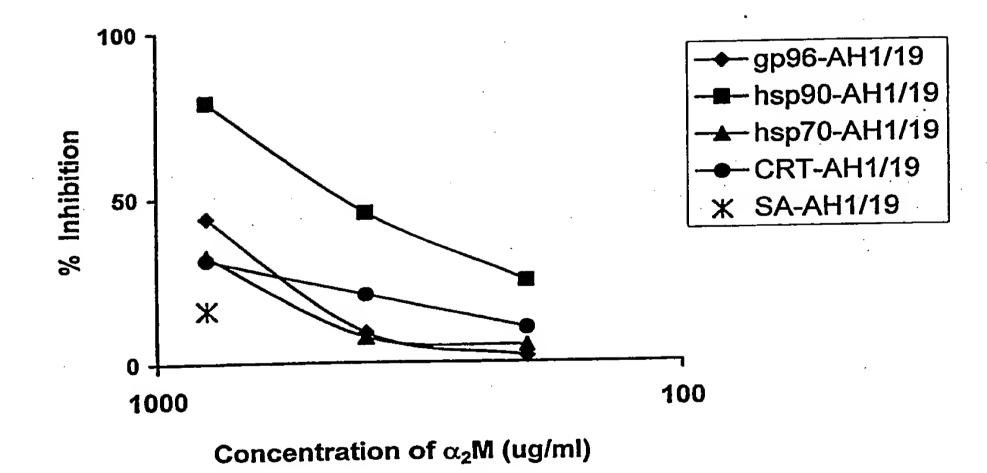
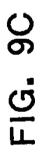
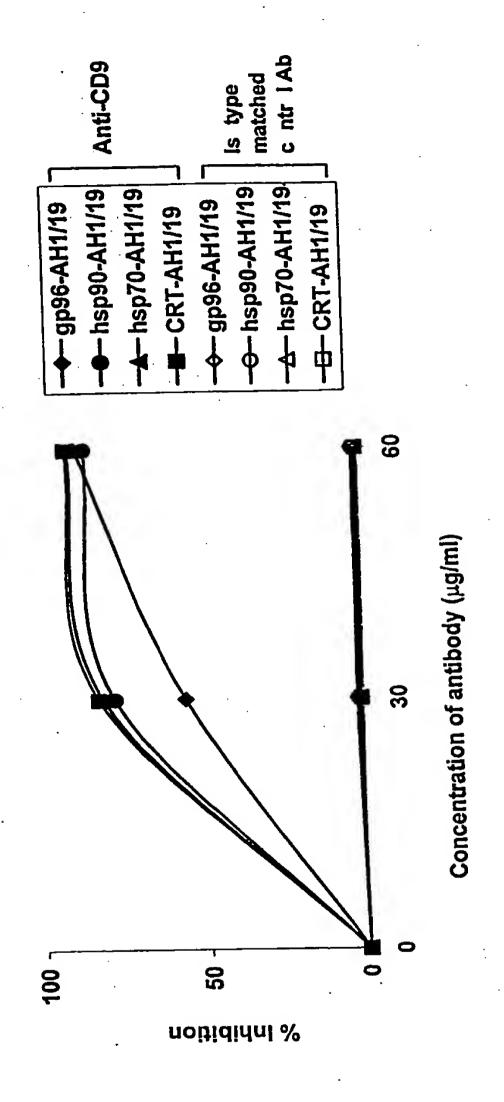


FIG. 9B





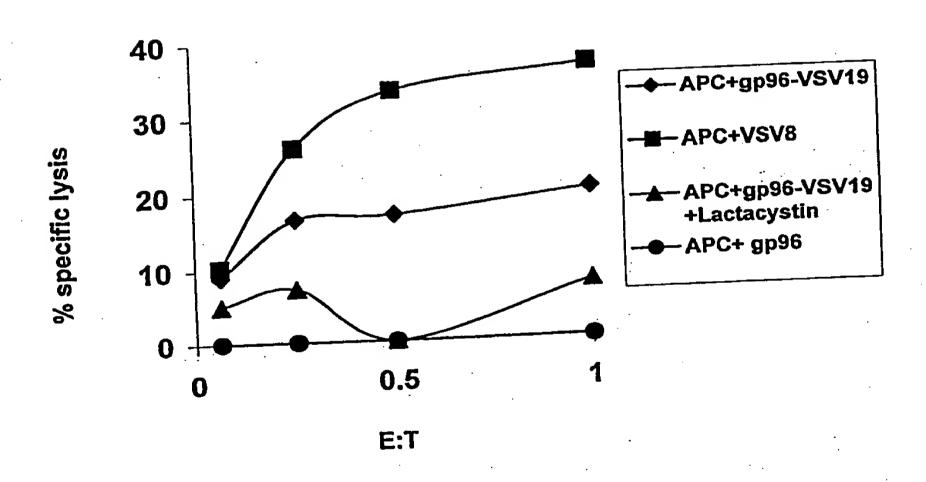


FIG. 10A

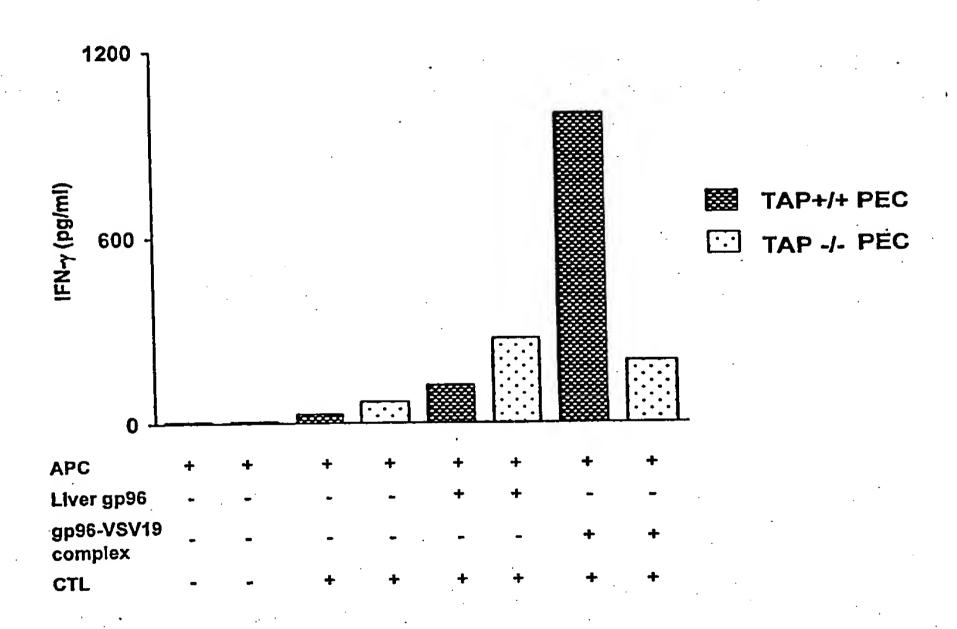


FIG. 10B

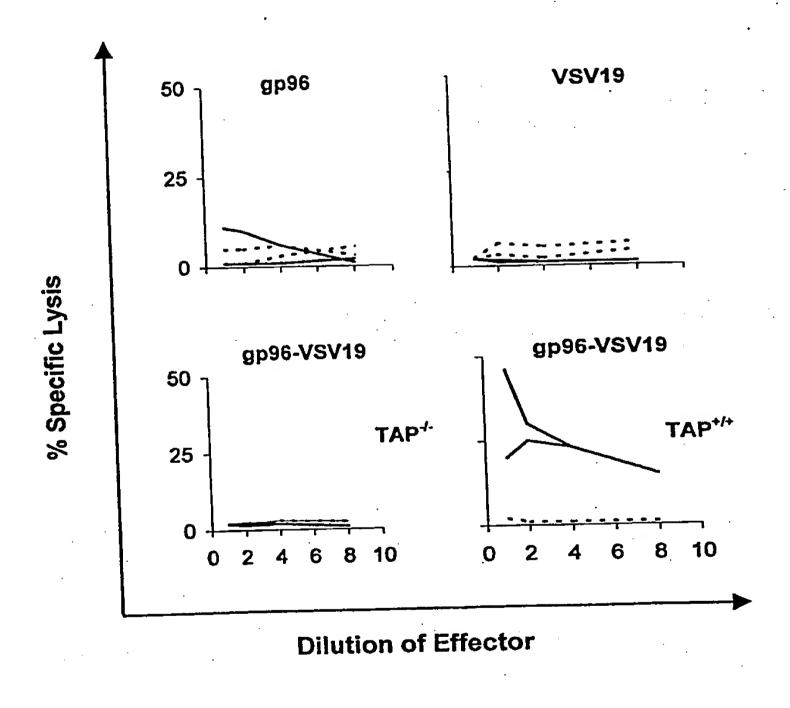


FIG. 10C

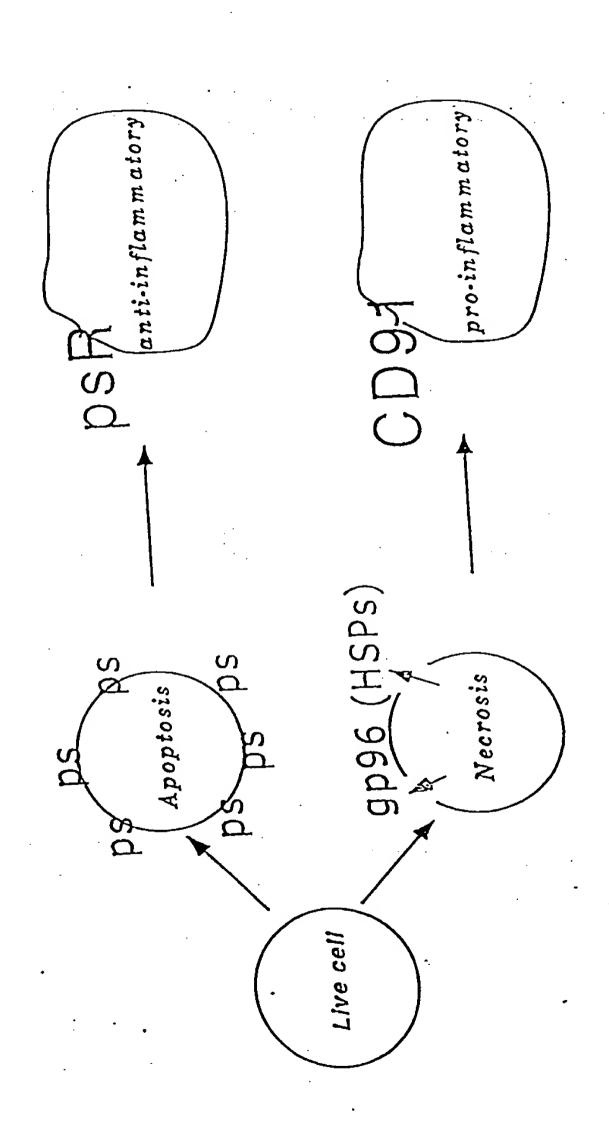


FIG. 11

GGCCC CAATT GAGGC CGCAC	CCTAC FGTGC FGGAC CCCGC FTTCC	C AF CA TI GA GO CG TO CC AI	AGGCA TTTTO AGCA CAGCA TTGCT	CCCC GCAGC AGGCC AGGCC TAAC	CAT CGG CTT GGA GGG CCF	CGGG AGTC AAGC CCCA AGGAT CGAGG	TCC CGGC CAGG CAGG CAAG CACA CTG	ACGC TCCG GGTG GGCT ATAG	CCCC BAGAT BAGG BAGG BAGTAI CCG (CCA (CECA) CECA (C	CCCC GGCTC GGCTA GGGCC GGACC GGACC	CCACO STGAC ATTTC CCATT SAGAC CAGAC CTG	CC CC SC T' SG GC TT CA SG AA CTG C	GCCTC TCGCC GGCA(ACCT) AGAT, TGGG CTC	CATCA CCTCC CCTGG GGGGG ATGCC AAAGG GGCTG GTG Val	60 120 180 240 300 360 420
CCG (CTG (Leu]	CTT :	Ser A	GCT (Ala 1 15	CTG (Leu \	STC 7	rcc (Ser (GGG (SCC A Ala 1 20	ACT . Thr	ATG (Met)	GAT (Asp	GCC Ala	CCT Pro 25	AAA Lys	519
ACT Thr	TGC : Cys :	Ser	CCT : Pro :	AAG (Lys (CAG (Gln)	rrr (Phe	GCC Ala	TGC I Cys I 35	AGA (Arg)	GAC Asp	CAA Glņ	Ile	ACC Thr 40	TGT Cys	ATC Ile	567
TCA Ser	AAG Lys	GGC Gly 45	TGG Trp	CGG '	TGT (Cys)	Asp (GGT Gly 50	GAA . Glu .	AGA Arg	GAT Asp	TGC Cys	CCC Pro 55	GAC Asp	GGC Gly	TCT Ser	615
GAT Asp	GAA Glu 60	GCC Ala	CCT Pro	GAG Glu	ATC Ile	TGT Cys 65	CCA Pro	CAG Gln	AGT Ser	AAA Lys	GCC Ala 70	CAG Gln	AGA Arg	TGC Cys	CCG Pro	663
CCA Pro 75	AAT Asn	GAG Glu	CAC His	AGT Ser	TGT Cys 80	CTG Leu	G1 y	ACT Thr	GAG Glu	CTA Leu 85	TGT Cys	GTC Val	CCC Pro	ATG Met	TCT Ser 90	711
CGT Arg	CTC Leu	TGC Cys	AAC Asn	GGG Gly 95	ATC Ile	CAG Gln	GAC Asp	TGC Cys	ATG Met 100	GAT Asp	GGC GGC	TCA Ser	GAC Asp	GAG Glu 105	Gry	759
GCT Ala	CAC His	TGC Cys	CGA Arg 110	GAG Glu	CTC Leu	CGA Arg	GCC Ala	AAC Asn 115	TGT Cys	TCT Ser	CGA Arg	ATG Met	GGT Gly 120	Cys	CAA Gln	807 .
CAC His	CAT His	TGT Cys 125	Val	CCT Pro	ACA Thr	CCC Pro	AGT Ser 130	Gly	CCC Pro	ACG Thr	TGC Cys	TAC Tyr 135	TGT Cys	AAC Asn	AGC Ser	855
AGC Ser	TTC Phe	Gln	CTC Leu	GAG Glu	GCA Ala	GAT Asp 145	Gly	AAG Lys	ACG Thr	TGC Cys	Lys 150	Asp	TTT Phe	GAC	GAG Glu	903
TG1 Cys 155	Ser	GTG Val	TAT Tyr	GGC	ACC Thr 160	Cys	AGC Ser	CAG Gln	CTT Leu	TGC Cys 165	Thr	CAA C	ACA Thi	A GA?	GGC Gly 170	951
TCC Sea	TTC Phe	ACA Thr	TG1	GGC Gly	Cys	GTT Val	GAI Glu	A GGC	TAC Tyr 180	: Le	CTC	G CAF	A CCC	G GAG o Asy 18	C AAC p Asn 5	999
CG(Ar	C TCC g Sei	TGC Cy:	C AA(s Ly: 19(s Ala	AAC Lys	FAA 3 Ra 3	GA(G CCF u Pro 19!	va!	A GA' L As	T CGO	g CC	G CC b Pr 20	o va	G CTA	1047

CTG I	[le	GCC Ala 205	AAC Asn	TCT Ser	CAG Gln	AAC Asn	ATC Ile 210	CTA Leu	GCT Ala	ACG Thr	TAC Tyr	CTG Leu 215	AGT Ser	GGG Gly	GCC Ala	1095
CAA (Gln '	GTG Val 220	TCT Ser	ACC Thr	ATC Ile	ACA Thr	CCC Pro 225	ACC Thr	AGC Ser	ACC Thr	CGA Arg	CAA Gln 230	ACC Thr	ACG Thr	GCC Ala	ATG Met	1143
GAC Asp 235	TTC Phe	AGT Ser	TAT Tyr	GCC Ala	AAT Asn 240	GIU	ACC Thr	GTA Val	TGC Cys	TGG Trp 245	Val	CAC	GTT Val	GT A	GAC Asp 250	1191
AGT Ser	GCT Ala	GCC Ala	CAG Gln	ACA Thr 255	Gln	CTC Leu	AAG Lys	TGT Cys	GCC Ala 260	Arg	ATG Met	CCT Pro	GGC	CTG Leu 265	232	1239
GGC Gly	TTT Phe	GTG Val	GAT Asp 270	GAG Glu	His	ACC	ATC Ile	AAC Asn 275	TIE	TCC Ser	CTC	AGC Ser	CTG Leu 280	1120	CAC His	1287
GTG Val	GAG Glu	CAG Glr 285	n Met	GCF Ala	ATC	GAC Asp	TGG Trp 290	rec	ACC Thi	GG#	AA A sa y	TTC Phe 295	: IYI	TTT Phe	GTC Val	1335
GAC Asp	GAC Asp	, Ile	r GAG 2 Ası	C GAG	C AGO	OTA 3	s Pue	r GTC e Val	TG:	AA 1 s Ası	C CG/ n Arc 310	y Asi	GGG Gly	GAC / Asp	ACC Thr	1383
TGT Cys 315	GT(T CT	u Le	u Asj	C CTO p Lev	i eri	n re	ату	I AS	U. ET	C AAI	A GGC s Gly	y Il	C GCC e Ala 330	1431
		C CC p Pr	C GC	C AT a Me 33	t GI	G AAG y Ly	G GT s Va	G TT	C TT e Ph 34	6 111	T GA r As	C TA	C GGC r Gl	G CA y Gl 34	G ATC n Ile 5	1479
CC# Pro	A AA	G GT s Va	G GA 1 G1 35	u Ar	C TG	T GA	C AT p Me	G GA t As	b er	A CA y Gl	AA D. eA n.	c CG	C AC g Th 36		G CTG	1527 •
GT(Va	G GA l As	T AC p Se 36	er Ly	AG AT (S I)	C GI Le Va	G TT	T CC ie Pr 37	OH	C GC Ls G	SC AT	C AC Le Tì	CC CI		C CI	G GTC eu Val	1575
AG Se	r Ai	GC C	rc G' eu V	rc TA	AC TO	cb v	CG G/ La A: 35	AC GC	CC TA	AC C	eu A	AC TA sp Ty 90	AC AT	rc G/ Le G/	AG GTG lu Val	1623
GT Va 39	l A	AC T.	AC G yr G	AA G lu G	\mathbf{r}	AG GG ys Gi	GT Co ly A	GG C. rg G	AG A ln T	UL I	TC A le I 05	TC C. le G	AA G	GC A' ly I	TC CTG le Leu 410	1671
		AG C lu H	AC C	eu T	AC G yr G 15	GC C ly L	TG Ā eu T	CC G	ar F	TT G he G	AG A Slu A	AC T sn T	AT C yr L		AC GCC yr Ala 25	1719
A(Ti	CC A	AC T	Ser F	AC A Asp A	AT G	CC A	AC A Isn I	inr (AG (61n (CAG A	AAG A Lys 1	ACG A Thr S		TG F al 1	ATC CGA [le Arg	1767

FIG. 12A

(Sheet 23 of 91)

0 7 70 10 1	•
GTG AAC CGG TTC AAC AGT ACT GAG TAC CAG GTC GTC ACC CGT GTO GTO GTC ACC CGT GTO GTO GTC ACC CGT GTO GTO GTO GTO GTO GTO GTO GTO GTO G	15
AAG GGT GCC CTG CAT ATC TAC CAC CAG CGC CAG CCC CON STATE AAG GGT GGT GCC CAG CCC CAG CAG	363
CGG AGT CAC GCC TGT GAG AAT GAC CAG TAC GGG AAG CCA GGT GGC TGC Arg Ser His Ala Cys Glu Asn Asp Gln Tyr Gly Lys Pro Gly Gly Cys 480 480	911
TCC GAC ATC TGC CTC CTG GCC AAC AGT CAC AAG GCA AGG ACC 1GC 1.50 Ser Asp Ile Cys Leu Leu Ala Asn Ser His Lys Ala Arg Thr Cys Arg 505	959
TGC AGG TCT GGC TTC AGC CTG GGA AGT GAT GGG AAG TCT TGT AAG AAA ZCVs Arg Ser Gly Phe Ser Leu Gly Ser Asp Gly Lys Ser Cys Lys 515	2007
CCT GAA CAT GAG CTG TTC CTC GTG TAT GGC AAG GGC CGA CCA GGC ATC Pro Glu His Glu Leu Phe Leu Val Tyr Gly Lys Gly Arg Pro Gly Ile 530 535	2055
ATT AGA GGC ATG GAC ATG GGG GCC AAG GTC CCA GAT GAG CAC ATG ATC Ile Arg Gly Met Asp Met Gly Ala Lys Val Pro Asp Glu His Met Ile 545	2103
CCC ATC GAG AAC CTT ATG AAT CCA CGC GCT CTG GAC TTC CAC GCC GAG Pro Ile Glu Asn Leu Met Asn Pro Arg Ala Leu Asp Phe His Ala Glu 560 565	2151
ACC GGC TTC ATC TAC TTT GCT GAC ACC ACC AGC TAC CTC ATT GGC CGC Thr Gly Phe Ile Tyr Phe Ala Asp Thr Thr Ser Tyr Leu Ile Gly Arg 585	2199
CAG AAA ATT GAT GGC ACG GAG AGA GAG ACT ATC CTG AAG GAT GGC ATC	2247 .
CAC AAT GTG GAG GGC GTA GCC GTG GAC TGG ATG GGA GAC AAT CTT TAC His Asn Val Glu Gly Val Ala Val Asp Trp Met Gly Asp Asn Leu Tyr 610 615	2295
TGG ACT GAT GGC CCC AAG AAG ACC ATT AGT GTG GCC AGG CTG GAG Trp Thr Asp Asp Gly Pro Lys Lys Thr Ile Ser Val Ala Arg Leu Glu 625	2343
AAA GCC GCT CAG ACC CGG AAG ACT CTA ATT GAG GGC AAG ATG ACA CAC Lys Ala Ala Gln Thr Arg Lys Thr Leu Ile Glu Gly Lys Met Thr His 650	2391
CCC AGG GCC ATT GTA GTG GAT CCA CTC AAT GGG TGG ATG TAC TGG ACA CCC AGG GCC ATT GTA GTG GAT CCA CTC AAT GGG TGG ATG TAC TGG ACA CCC AGG GCC ATT GTA GTG GAT CCA CTC AAT GGG TGG ATG TAC TGG ACA CCC AGG GCC ATT GTA GTG GAT CCA CTC AAT GGG TGG ATG TAC TGG ACA CCC AGG GCC ATT GTA GTG GAT CCA CTC AAT GGG TGG ATG TAC TGG ACA CCC AGG GCC ATT GTA GTG GAT CCA CTC AAT GGG TGG ATG TAC TGG ACA CCC AGG GCC ATT GTA GTG GAT CCA CTC AAT GGG TGG ATG TAC TGG ACA CCC AGG GCC ATT GTA GTG GAT CCA CTC AAT GGG TGG ATG TAC TGG ACA CCC AGG GCC ATT GTA GTG GAT CCA CTC AAT GGG TGG ATG TAC TGG ACA CCC AGG GCC ATT GTA GTG GAT CCA CTC AAT GGG TGG ATG TAC TGG ACA CCC AGG GCC ATT GTA GTG GAT CCA CTC AAT GGG TGG ATG TAC TGG ACA CCC AGG GCC ATT GTA GTG GAT CCA CTC AAT GGG TGG ATG TAC TGG ACA CCC AGG GCC ATT GTA GTG GAT CCA CTC AAT GGG TGG ATG TAC TGG ACA CCC AGG GCC ATT GTA GTG GAT CCA CTC AAT GGG TGG ATG TAC TGG ACA CCC AGG GCC ATT GTA GTG GAT CCA CTC AAT GGG TGG ATG TAC TGG ACA CCC AGG GCC ATT GTA GTG AGG ATG TAC TGG ATG TGG ATG TAC TGG ACA CCC AGG ATG ALG ALG TGG ATG TAC TGG ATG TGG AT	2439
GAC TGG GAG GAC CCC AAG GAC AGT CGG CGA GGG CGG CTC GAG AGG ASD Trp Glu Glu Asp Pro Lys Asp Ser Arg Arg Gly Arg Leu Glu Arg	2487
670	

(Sheet 24 of 91)

· •	
GCT TGG ATG GAC GGC TCA CAC CGA GAT ATC TTT GTC ACC TCC AAG ACA Ala Trp Met Asp Gly Ser His Arg Asp Ile Phe Val Thr Ser Lys Thr 690 695	535
GTG CTT TGG CCC AAT GGG CTA AGC CTG GAT ATC CCA GCC GGA CGC CTC 2 Val Leu Trp Pro Asn Gly Leu Ser Leu Asp Ile Pro Ala Gly Arg Leu 705	583
TAC TGG GTG GAT GCC TTC TAT GAC CGA ATT GAG ACC ATA CIG GTO IAST Tyr Trp Val Asp Ala Phe Tyr Asp Arg Ile Glu Thr Ile Leu Leu Asn 720 725	631
	267 _. 9
TTC GGC CTG TGT CAC CAT GGC AAC TAC CTC TTT TGG ACC GAG TAC CGG Phe Gly Leu Cys His His Gly Asn Tyr Leu Phe Trp Thr Glu Tyr Arg 750 750	2727
AGC GGC AGC GTC TAC CGC TTG GAA CGG GGC GTG GCA GGC GCA CCG CCC Ser Gly Ser Val Tyr Arg Leu Glu Arg Gly Val Ala Gly Ala Pro Pro	2775
ACT GTG ACC CTT CTG CGC AGC GAG AGA CCG CCT ATC TTT GAG ATC CGA Thr Val Thr Leu Leu Arg Ser Glu Arg Pro Pro Ile Phe Glu Ile Arg 785	2823
ATG TAC GAC GCG CAC GAG CAG CAA GTG GGT ACC AAC AAA TGC CGG GTA Met Tyr Asp Ala His Glu Gln Gln Val Gly Thr Asn Lys Cys Arg Val 800 805	2871
AAT AAC GGA GGC TGC AGC AGC CTG TGC CTC GCC ACC CCC GGG AGC CGC AAT AAC GGA GGC TGC AGC AGC CTG TGC CTC GCC ACC CCC GGG AGC CGC AAT AAC GGA GGC TGC AGC CTG TGC CTC GCC ACC CCC GGG AGC CGC AAT AAC GGA GGC TGC AGC CTG TGC CTC GCC ACC CCC GGG AGC CGC AAT AAC GGA GGC TGC AGC CTG TGC CTC GCC ACC CCC GGG AGC CGC AAT AAC GGA GGC TGC AGC CTG TGC CTC GCC ACC CCC GGG AGC CGC AAT AAC GGA GGC TGC AGC AGC CTG TGC CTC GCC ACC CCC GGG AGC CGC AAT AAC GGA GGC TGC AGC AGC CTG TGC CTC GCC ACC CCC GGG AGC CGC AAT AAC GGA GGC TGC AGC AGC CTG TGC CTC GCC ACC CCC GGG AGC CGC AAT AAC GGA GGC TGC AGC AGC CTG TGC ACC CCC GGG AGC CGC AAT AAC GGA GGC TGC AGC AGC CTG TGC ACC CCC GGG AGC CGC AAT AAC GGA GGC TGC AGC AGC CTG TGC ACC CCC GGG AGC CGC AAT AAC GGA GGC TGC AGC AGC CTG TGC ACC CCC GGG AGC CGC AAT AAC GGA GGC TGC AGC AGC AGC CTG TGC ACC CCC GGG AGC CGC AAT AAC GGA GGC TGC AGC AGC AGC CTG TGC ACC ACC CCC GGG AGC CGC AAT AAC GGA GGC TGC AGC AGC AGC AGC CTG TGC ACC ACC CCC GGG AGC CGC AAT AAC GGA GGC TGC AGC AGC AGC CTG TGC ACC ACC ACC ACC ACC ACC ACC ACC ACC A	2919
CAG TGT GCC TGT GCC GAG GAC CAG GTG TTG GAC ACA GAT GGT GTC ACC Gln Cys Ala Cys Ala Glu Asp Gln Val Leu Asp Thr Asp Gly Val Thr	2967
TGC TTG GCG AAC CCA TCC TAC GTG CCC CCA CCC CAG TGC CAG CCG GGC TGC TTG GCG AAC CCA TCC TAC GTG CCC CCA CCC CAG TGC CAG CCG GGC TGC TTG GCG AAC CCA TCC TAC GTG CCC CCA CCC CAG TGC CAG CCG GGC TGC TTG GCG AAC CCA TCC TAC GTG CCC CCA CCC CAG TGC CAG CCG GGC TGC TTG GCG AAC CCA TCC TAC GTG CCC CCA CCC CAG TGC CAG CCG GGC TGC TTG GCG AAC CCA TCC TAC GTG CCC CCA CCC CAG TGC CAG CCG GGC TGC TTG GCG AAC CCA TCC TAC GTG CCC CCA CCC CAG TGC CAG CCG GGC TGC TTG GCG AAC CCA TCC TAC GTG CCC CCA CCC CAG TGC CAG CCG GGC TGC TTG GCG AAC CCA TCC TAC GTG CCC CCA CCC CAG TGC CAG CCG GGC TGC TTG GCG AAC CCA TCC TAC GTG CCC CCA CCC CAG TGC CAG CCG GGC TGC TTG GCG AAC CCA TCC TAC GTG CCC CCA CCC CAG TGC CAG CCG GGC TGC TTG GCG AAC CCA TCC TAC GTG CCC CCA CCC CAG TGC CAG CCG GGC TGC TTG GCG AAC CCA TCC TAC GTG CCC CCA CCC CAG TGC CAG CCG GGC TGC TTG GCG AAC CCA TCC TAC GTG CCC CCA CCC CAG TGC CAG CCG GGC TGC TTG TTG TTG TTG TTG TTG TTG TTG TTG	3015
CAG TTT GCC TGT GCC AAC AAC CGC TGC ATC CAG GAG CGC TGG AAG TGT Gln Phe Ala Cys Ala Asn Asn Arg Cys Ile Gln Glu Arg Trp Lys Cys 865	3063
GAC GGA GAC AAC GAC TGT CTG GAC AAC AGC GAT GAG GCC CCA GCA CTG Asp Gly Asp Asn Asp Cys Leu Asp Asn Ser Asp Glu Ala Pro Ala Leu 890	3111
TGC CAT CAA CAC ACC TGT CCC TCG GAC CGA TTC AAG TGT GAG AAC AAC TGC CAT CAA CAC ACC TGT CCC TCG GAC CGA TTC AAG TGT GAG AAC AAC TGC CAT CAA CAC ACC TGT CCC TCG GAC CGA TTC AAG TGT GAG AAC AAC TGC CAT CAA CAC ACC TGT CCC TCG GAC CGA TTC AAG TGT GAG AAC AAC TGC CAT CAA CAC ACC TGT CCC TCG GAC CGA TTC AAG TGT GAG AAC AAC TGC CAT CAA CAC ACC TGT CCC TCG GAC CGA TTC AAG TGT GAG AAC AAC TGC CAT CAA CAC ACC TGT CCC TCG GAC CGA TTC AAG TGT GAG AAC AAC TGC CAT CAA CAC ACC TGT CCC TCG GAC CGA TTC AAG TGT GAG AAC AAC TGC CAT CAA CAC ACC TGT CCC TCG GAC CGA TTC AAG TGT GAG AAC AAC TGC CAT CAA CAC ACC TGT CCC TCG GAC CGA TTC AAG TGT GAG AAC AAC TGC CAT CAA CAC ACC TGT CCC TCG GAC CGA TTC AAG TGT GAG AAC AAC TGC CAT CAA CAC ACC TGT CCC TCG GAC CGA TTC AAG TGT GAG AAC AAC TGC CAT CAA CAC ACC TGT CCC TCG GAC CGA TTC AAG TGT GAG AAC AAC TGC CAT CAA CAC ACC TGT CCC TCG GAC CGA TTC AAG TGT GAG AAC AAC TGC CAT CAA CAC ACC TGT CCC TCG GAC CGA TTC AAG TGT GAG AAC AAC TGC CAT TGT TGT TGT TGT TGT TGT TGT TGT TGT T	3159
CGG TGT ATC CCC AAC CGC TGG CTC TGT GAT GGG GAT AAT GAT TGT GGC Arg Cys Ile Pro Asn Arg Trp Leu Cys Asp Gly Asp Asn Asp Cys Gly 910 915	3207

(Sheet 25 of 91)

													ACC Thr			3255
										Cys			ATC Ile			3303
ACC Thr 955	TGT Cys	GAT Asp	CTG Leu	GAT Asp	GAT Asp 960	GAC Asp	TGT Cys	GJ Y GGG	GAC Asp	CGG Arg 965	TCC Ser	GAT Asp	GAG Glu	TCA Ser	GCC Ala 970	3351
TCA Ser	TGC Cys	GCC Ala	TAC Tyr	CCC Pro 975	ACC Thr	TGC Cys	TTC Phe	CCC Pro	CTG Leu 980	ACT Thr	CAA Gln	TTT Phe	ACC Thr	TGC Cys 985	AAC Asn	3399
AAT Asn	GGC Gly	AGA Arg	TGT Cys 990	ATT Ile	AAC Asn	ATC Ile	AAC Asn	TGG Trp 995	CGG Arg	TGT Cys	GAC Asp	Asn	GAC Asp 1000	AAT Asn	GAC Asp	3447
TGT Cys	Gly	GAC Asp 1005	AAC Asn	AGC Ser	GAC Asp	Glu	GCC Ala 1010	GGC Gly	TGC Cys	AGT Ser	His	TCC Ser 1015	TGC Cys	TCC Ser	AGT Ser	3495
Thr	CAG Gln 1020	TTC Phe	AAG Lys	TGC Cys	Asn	AGT Ser 1025	GGC Gly	AGA Arg	TGC Cys	Ile	CCC Pro 1030	GAG Glu	CAC His	TGG Trp	ACG Thr	3543
TGT Cys 1035	GAT Asp	GGG Gly	GAC Asp	Asn	GAT Asp 1040	Cys	GGG Gly	GAC Asp	Tyr	AGC Ser 1045	Asp	GAG Glu	ACA Thr	His	GCC Ala 1050	3591
AAC Asn	TGT Cys	ACC Thr	Asn	CAG Gln 1055	Ala	ACA Thr	AGA Arg	Pro	CCT Pro 1060	Gly	Gly	TGC Cys	CAC His	TCG Ser 1065	Asp	3639
GAG Glu	TTC Phe	CAG Gln	TGC Cys 1070	Pro	CTA Leu	GAT Asp	Gly	CTG Leu 1075	Cys	ATC	Pro	CTG Leu	AGG Arg 1080	Trp	CGC Arg	3687
TGC Cys	Asp	GGG Gly 1085	Asp	ACC Thr	GAC Asp	TGC Cys	ATG Met 1090	Asp	TCC Ser	AGC Ser	GAT Asp	GAG Glu 1095	ı Lys	AGC Ser	TGT Cys	3735
GAG Glu	GGC Gly 1100	· Val	ACC Thr	CAT His	: Val	TGT Cys 1105	Asp	CCG Pro	TAA 3 neA c	GTC Val	: AAG . Lys 1110	; Ph∈	GG GG GG GG GG GG GG GG GG GG GG GG GG	TGC Cys	AAG Lys	3783
GAC Asp 1115	Ser	GCC Ala	C CGC	TGC Cys	ATC 1120	e Ser	AAC Lys	G GCC	TGG Trp	GTG Val 1125	Cys	GAT S Asp	GGC GL	C GAC	Ser 1130	3831
GAC Asp	TG7 Cys	GA Glu	A GA?	7 AAG 2 Ast 113	ı Se	C GAC	GAC Glu	i GJ	3 AA(u As: 114(n Cys	GA(G GCG	C CT(3 GCC u Ala 114	TGC TGC A Cys	3879
AG(G CC/	A CC	C TC o Se: 115	r His	r CCC	C TG(o Cy:	C GCG s Ala	C AAG a Ass 115	n Ası	n Th	C TC	T GT r Va	C TG 1 Cy 116	s Le	G CCT u Pro	3927

FIG. 12A

8449-134 (Sheet 26 of 91)

CCT GAC AAG CTG TGC GAC GGC AAG GAT GAC TGT GGA GAC GGC TCG GAT Pro Asp Lys Leu Cys Asp Gly Lys Asp Asp Cys Gly Asp Gly Ser Asp 1165 1170 1175	3975
GAG GGC GAG CTC TGT GAC CAG TGT TCT CTG AAT AAT GGT GGC TGT AGT Glu Gly Glu Leu Cys Asp Gln Cys Ser Leu Asn Asn Gly Gly Cys Ser 1180	4023
CAC AAC TGC TCA GTG GCC CCT GGT GAA GGC ATC GTG TGC TCT TGC CCT His Asn Cys Ser Val Ala Pro Gly Glu Gly Ile Val Cys Ser Cys Pro 1200 1205	4071
CTG GGC ATG GAG CTG GGC TCT GAC AAC CAC ACC TGC CAG ATC CAG AGC Leu Gly Met Glu Leu Gly Ser Asp Asn His Thr Cys Gln Ile Gln Ser 1215 1220 1225	4119
TAC TGT GCC AAG CAC CTC AAA TGC AGC CAG AAG TGT GAC CAG AAC AAG Tyr Cys Ala Lys His Leu Lys Cys Ser Gln Lys Cys Asp Gln Asn Lys 1230 1235	4167
TTC AGT GTG AAG TGC TCC TGC TAC GAG GGC TGG GTC TTG GAG CCT GAC Phe Ser Val Lys Cys Ser Cys Tyr Glu Gly Trp Val Leu Glu Pro Asp 1245 1250 1255	4215
GGG GAA ACG TGC CGC AGT CTG GAT CCC TTC AAA CTG TTC ATC ATC TTC Gly Glu Thr Cys Arg Ser Leu Asp Pro Phe Lys Leu Phe Ile Ile Phe 1260 1265 1270	4263
TCC AAC CGC CAC GAG ATC AGG CGC ATT GAC CTT CAC AAG GGG GAC TAC Ser Asn Arg His Glu Ile Arg Arg Ile Asp Leu His Lys Gly Asp Tyr 1275 1280 1285	4311
AGC GTC CTA GTG CCT GGC CTG CGC AAC ACT ATT GCC CTG GAC TTC CAC Ser Val Leu Val Pro Gly Leu Arg Asn Thr Ile Ala Leu Asp Phe His 1295 1300 1305	4359
CTC AGC CAG AGT GCC CTC TAC TGG ACC GAC GCG GTA GAG GAC AAG ATC Leu Ser Gln Ser Ala Leu Tyr Trp Thr Asp Ala Val Glu Asp Lys Ile 1310	4407
TAC CGT GGG AAA CTC CTG GAC AAC GGA GCC CTG ACC AGC TTT GAG GTG Tyr Arg Gly Lys Leu Leu Asp Asn Gly Ala Leu Thr Ser Phe Glu Val 1325 1330 1335	4455
GTG ATT CAG TAT GGC TTG GCC ACA CCA GAG GGC CTG GCT GTA GAT TGG Val Ile Gln Tyr Gly Leu Ala Thr Pro Glu Gly Leu Ala Val Asp Trp 1340 1345 1350	4503
ATT GCA GGC AAC ATC TAC TGG GTG GAG AGC AAC CTG GAC CAG ATC GAA Ile Ala Gly Asn Ile Tyr Trp Val Glu Ser Asn Leu Asp Gln Ile Glu 1355 1360 1365	4551
GTG GCC AAG CTG GAC GGA ACC CTC CGA ACC ACT CTG CTG GCG GGT GAC Val Ala Lys Leu Asp Gly Thr Leu Arg Thr Thr Leu Leu Ala Gly Asp 1375	4599
ATT GAG CAC CCG AGG GCC ATC GCT CTG GAC CCT CGG GAT GGG ATT CTG Ile Glu His Pro Arg Ala Ile Ala Leu Asp Pro Arg Asp Gly Ile Leu 1390 1395 1400	4647

FIG. 12A

TTT TGG ACA GAC TGG GAT GCC AGC CTG CCA CGA ATC GAG GCT GCA TCC Phe Trp Thr Asp Trp Asp Ala Ser Leu Pro Arg Ile Glu Ala Ala Ser 1405 1410 1415	4695
ATG AGT GGA GCT GGC CGC CGA ACC ATC CAC CGG GAG ACA GGC TCT GGG Met Ser Gly Ala Gly Arg Arg Thr Ile His Arg Glu Thr Gly Ser Gly 1420	4743
GGC TGC GCC AAT GGG CTC ACC GTG GAT TAC CTG GAG AAG CGC ATC CTC Gly Cys Ala Asn Gly Leu Thr Val Asp Tyr Leu Glu Lys Arg Ile Leu 1435	4791
TGG ATT GAT GCT AGG TCA GAT GCC ATC TAT TCA GCC CGG TAT GAC GGC Trp Ile Asp Ala Arg Ser Asp Ala Ile Tyr Ser Ala Arg Tyr Asp Gly 1455 1460 1465	4839
TCC GGC CAC ATG GAG GTG CTT CGG GGA CAC GAG TTC CTG TCA CAC CCA Ser Gly His Met Glu Val Leu Arg Gly His Glu Phe Leu Ser His Pro 1470 1475	4887
TTT GCC GTG ACA CTG TAC GGT GGG GAG GTG TAC TGG ACC GAC TGG CGA Phe Ala Val Thr Leu Tyr Gly Glu Val Tyr Trp Thr Asp Trp Arg 1495	4935
ACA AAT ACA CTG GCT AAG GCC AAC AAG TGG ACT GGC CAC AAC GTC ACC Thr Asn Thr Leu Ala Lys Ala Asn Lys Trp Thr Gly His Asn Val Thr 1500 1505	4983
GTG GTA CAG AGG ACC AAC ACC CAG CCC TTC GAC CTG CAG GTG TAT CAC Val Val Gln Arg Thr Asn Thr Gln Pro Phe Asp Leu Gln Val Tyr His 1515	5031
CCT TCC CGG CAG CCC ATG GCT CCA AAC CCA TGT GAG GCC AAT GGC GGC Pro Ser Arg Gln Pro Met Ala Pro Asn Pro Cys Glu Ala Asn Gly Gly 1535	5079
CGG GGC CCC TGT TCC CAT CTG TGC CTC ATC AAC TAC AAC CGG ACC GTC Arg Gly Pro Cys Ser His Leu Cys Leu Ile Asn Tyr Asn Arg Thr Val 1550	5127 ·
TCC TGG GCC TGT CCC CAC CTC ATG AAG CTG CAC AAG GAC AAC ACC ACC Ser Trp Ala Cys Pro His Leu Met Lys Leu His Lys Asp Asn Thr Thr 1575	5175
TGC TAT GAG TTT AAG AAG TTC CTG CTG TAC GCA CGT CAG ATG GAG ATC Cys Tyr Glu Phe Lys Lys Phe Leu Leu Tyr Ala Arg Gln Met Glu Ile 1580 1590	5223
CGG GGC GTG GAC CTG GAT GCC CCG TAC TAC AAT TAT ATC ATC TCC TTC Arg Gly Val Asp Leu Asp Ala Pro Tyr Tyr Asn Tyr Ile Ile Ser Phe 1600 1605	5271 ·
ACG GTG CCT GAT ATC GAC AAT GTC ACG GTG CTG GAC TAT GAT GCC CGA Thr Val Pro Asp Ile Asp Asn Val Thr Val Leu Asp Tyr Asp Ala Arg 1615 1620 1625	5319
GAG CAG CGA GTT TAC TGG TCT GAT GTG CGG ACT CAA GCC ATC AAA AGG Glu Gln Arg Val Tyr Trp Ser Asp Val Arg Thr Gln Ala Ile Lys Arg 1630 1640	5367

FIG. 12A

	5415
GCA TTT ATC AAC GGC ACT GGC GTG GAG ACC GTT GTC TCT GCA GAC TTG Ala Phe Ile Asn Gly Thr Gly Val Glu Thr Val Val Ser Ala Asp Leu 1645 1650 1655	3.23
CCC AAC GCC CAC GGG CTG GCT GTG GAC TGG GTC TCC CGA AAT CTG TTT Pro Asn Ala His Gly Leu Ala Val Asp Trp Val Ser Arg Asn Leu Phe 1660 1665 1670	5463
TGG ACA AGT TAC GAC ACC AAC AAG AAG CAG ATT AAC GTG GCC CGG CTG Trp Thr Ser Tyr Asp Thr Asn Lys Lys Gln Ile Asn Val Ala Arg Leu 1675 1680 1685 1690	5511
GAC GGC TCC TTC AAG AAT GCG GTG GTG CAG GGC CTG GAG CAG CCC CAC Asp Gly Ser Phe Lys Asn Ala Val Val Gln Gly Leu Glu Gln Pro His 1695	5559
GGC CTG GTC GTC CAC CCG CTT CGT GGC AAG CTC TAC TGG ACT GAT GGG Gly Leu Val Val His Pro Leu Arg Gly Lys Leu Tyr Trp Thr Asp Gly 1710 1715	5607
GAC AAC ATC AGC ATG GCC AAC ATG GAT GGG AGC AAC CAC ACT CTG CTC Asp Asn Ile Ser Met Ala Asn Met Asp Gly Ser Asn His Thr Leu Leu 1725	5655
TTC AGT GGC CAG AAG GGC CCT GTG GGG TTG GCC ATT GAC TTC CCT GAG Phe Ser Gly Gln Lys Gly Pro Val Gly Leu Ala Ile Asp Phe Pro Glu 1740 1745 1750	5703
AGC AAA CTC TAC TGG ATC AGC TCT GGG AAC CAC ACA ATC AAC CGT TGC Ser Lys Leu Tyr Trp Ile Ser Ser Gly Asn His Thr Ile Asn Arg Cys 1755 1760 1765	5751
AAT CTG GAT GGG AGC GAG CTG GAG GTC ATC GAC ACC ATG CGG AGC CAG Asn Leu Asp Gly Ser Glu Leu Glu Val Ile Asp Thr Met Arg Ser Gln 1775 1780 1785	5799
CTG GGC AAG GCC ACT GCC CTG GCC ATC ATG GGG GAC AAG CTG TGG TGG Leu Gly Lys Ala Thr Ala Leu Ala Ile Met Gly Asp Lys Leu Trp Trp 1790 1795 1800	5847 .
GCA GAT CAG GTG TCA GAG AAG ATG GGC ACG TGC AAC AAA GCC GAT GGC Ala Asp Gln Val Ser Glu Lys Met Gly Thr Cys Asn Lys Ala Asp Gly 1805	5895
TCT GGG TCC GTG GTG CTG CGG AAC AGT ACC ACG TTG GTT ATG CAC ATG Ser Gly Ser Val Val Leu Arg Asn Ser Thr Thr Leu Val Met His Met 1820 1825 1830	5943
AAG GTG TAT GAC GAG AGC ATC CAG CTA GAG CAT GAG GGC ACC AAC CCC Lys Val Tyr Asp Glu Ser Ile Gln Leu Glu His Glu Gly Thr Asn Pro 1835 1840 1845 1850	5991
TGC AGT GTC AAC AAC GGA GAC TGT TCC CAG CTC TGC CTG CCA ACA TCA Cys Ser Val Asn Asn Gly Asp Cys Ser Gln Leu Cys Leu Pro Thr Ser 1855 1860 1865	6039
GAG ACG ACT CGC TCC TGT ATG TGT ACA GCC GGT TAC AGC CTC CGG AGC Glu Thr Thr Arg Ser Cys Met Cys Thr Ala Gly Tyr Ser Leu Arg Ser 1870	6087

GGA CAG CAG GCC TGT GAG GGT GTG GGC TCT TTT CTC CTG TAC TOTAL GGA CAG CAG GCC TGT GAG GGT GTG GGC TCT TTT CTC CTG TAC TOTAL GGA GGA CAG GCC TGT GAG GGC TCT TTT CTC CTG TAC TGT GAG GGC TCT TTT CTC CTG TGT GAG GGC TCT TTT CTC CTG TGT GAG GGC TCT TTT GAG GGC TTT TTT GAG GG	135
CAT GAG GGA ATT CGG GGG ATT CCA CTA GAT CCC AAT GAC AAG 100 DATE CAT GAG GGG ATT CCA CTA GAT CCC AAT GAC AAG 100 DATE CAT GAC AAC AAG 100 DATE CAT GAC AAC AAC AAC AAC AAC AAC AAC AAC AAC	183
GCC CTG GTC CCA GTG TCC GGA ACT TCA CTG GCT GTC GGA ATC GAC TTC Ala Leu Val Pro Val Ser Gly Thr Ser Leu Ala Val Gly Ile Asp Phe 1920 1925 1930	5231
CAT GCC GAA AAT GAC ACT ATT TAT TGG GTG GAT ATG GGC CTA AGC ACC CAT GCC GAA AAT GAC ACT ATT TAT TGG GTG GAT ATG GGC CTA AGC ACC His Ala Glu Asn Asp Thr Ile Tyr Trp Val Asp Met Gly Leu Ser Thr 1945	6279
ATC AGC AGG GCC AAG CGT GAC CAG ACA TGG CGA GAG GAT GTG GTG ACC Ile Ser Arg Ala Lys Arg Asp Gln Thr Trp Arg Glu Asp Val Val Thr 1950 1955	6327
AAC GGT ATT GGC CGT GTG GAG GGC ATC GCC GTG GAC TGG ATC GCA GGC Asn Gly Ile Gly Arg Val Glu Gly Ile Ala Val Asp Trp Ile Ala Gly 1975	6375
AAC ATA TAC TGG ACG GAC CAG GGC TTC GAT GTC ATC GAG GTT GCC CGG Asn Ile Tyr Trp Thr Asp Gln Gly Phe Asp Val Ile Glu Val Ala Arg 1985 1990	6423
CTC AAT GGC TCT TTT CGT TAT GTG GTC ATT TCC CAG GGT CTG GAC AAG CTC AAT GGC TCT TTT CGT TAT GTG GTC ATT TCC CAG GGT CTG GAC AAG CTC AAT GGC TCT TTT CGT TAT GTG GTC ATT TCC CAG GGT CTG GAC AAG CTC AAT GGC TCT TTT CGT TAT GTG GTC ATT TCC CAG GGT CTG GAC AAG CTC AAT GGC TCT TTT CGT TAT GTG GTC ATT TCC CAG GGT CTG GAC AAG CTC AAT GGC TCT TTT CGT TAT GTG GTC ATT TCC CAG GGT CTG GAC AAG CTC AAT GGC TCT TTT CGT TAT GTG GTC ATT TCC CAG GGT CTG GAC AAG CTC AAT GGC TCT TTT CGT TAT GTG GTC ATT TCC CAG GGT CTG GAC AAG Leu Asn Gly Ser Phe Arg Tyr Val Val Ile Ser Gln Gly Leu Asp Lys 2010	6471
CCT CGG GCC ATC ACT GTC CAC CCA GAG AAG GGG TAC TTG TTC TGG ACC Pro Arg Ala Ile Thr Val His Pro Glu Lys Gly Tyr Leu Phe Trp Thr 2025	6519
GAG TGG GGT CAT TAC CCA CGT ATT GAG CGG TCT CGC CTT GAT GGC ACA Glu Trp Gly His Tyr Pro Arg Ile Glu Arg Ser Arg Leu Asp Gly Thr 2030 2035 2040	6567
GAG AGA GTG GTG TTG GTT AAT GTC AGC ATC AGC TGG CCC AAT GGC ATC Glu Arg Val Val Leu Val Asn Val Ser Ile Ser Trp Pro Asn Gly Ile 2055	6615
TCA GTA GAC TAT CAG GGC GGC AAG CTC TAC TGG TGT GAT GCT CGG ATG TCA GTA GAC TAT CAG GGC GGC AAG CTC TAC TGG TGT GAT GCT CGG ATG TCA GTA GAC TAT CAG GGC GGC AAG CTC TAC TGG TGT GAT GCT CGG ATG TCA GTA GAC TAT CAG GGC GGC AAG CTC TAC TGG TGT GAT GCT CGG ATG TCA GTA GAC TAT CAG GGC GGC AAG CTC TAC TGG TGT GAT GCT CGG ATG TCA GTA GAC TAT CAG GGC GGC AAG CTC TAC TGG TGT GAT GCT CGG ATG TCA GTA GAC TAT CAG GGC GGC AAG CTC TAC TGG TGT GAT GCT CGG ATG TCA GTA GAC TAT CAG GGC GGC AAG CTC TAC TGG TGT GAT GCT CGG ATG TCA GTA GAC TAT CAG GGC GGC AAG CTC TAC TGG TGT GAT GCT CGG ATG TCA GTA GAC TAT CAG GGC GGC AAG CTC TAC TGG TGT GAT GCT CGG ATG TCA GTA GAC TAT CAG GGC GGC AAG CTC TAC TGG TGT GAT GCT CGG ATG TCA GTA GAC TAT CAG GGC GGC AAG CTC TAC TGG TGT GAT GCT CGG ATG TCA GTA GAC TAT CAG GGC GGC AAG CTC TAC TGG TGT GAT GCT CGG ATG TCA GTA GAC TAT CAG GGC GGC AAG CTC TAC TGG TGT CGG ATG TCA GTA GAC TAT CAG GGC GGC AAG CTC TAC TGG TGT GAT GCT CGG ATG TCA GTA GAC TAT CAG GGC GGC AAG CTC TAC TGG TGT GAT GCT CGG ATG TCA GTA GAC TAT CAG GGC GGC AAG CTC TAC TGG TGT GAT GCT CGG ATG TCA GTA GAC TAT CAG GGC GGC AAG CTC TAC TGG TGT GAT GCT CGG ATG TCA GTA GAC TAT GAC TGG TGT GAT GCT CGG ATG	6663
GAC AAG ATC GAG CGC ATC GAC CTG GAA ACG GGC GAG AAC CGG GAG GTG Asp Lys Ile Glu Arg Ile Asp Leu Glu Thr Gly Glu Asn Arg Glu Val 2080 2085	6711
GTC CTG TCC AGC AAT AAC ATG GAT ATG TTC TCC GTG TCC GTG TTT GAG Val Leu Ser Ser Asn Asn Met Asp Met Phe Ser Val Ser Val Phe Glu 2105	6759
2095 GAC TTC ATC TGG AGT GAC AGA ACT CAC GCC AAT GGC TCC ATC AAG Asp Phe Ile Tyr Trp Ser Asp Arg Thr His Ala Asn Gly Ser Ile Lys 2110 2110	6807

(She t 30 of 91)

CGC Arg	Gly	TGC Cys 125	AAA Lys	GAC . Asp .	AAT (Asn <i>l</i>	SCT AG Ala Ti 21	hr A	AC 7	rcc (Ser 1	GTG (Val 1	Pro	CTG . Leu . 135	AGG Arg	ACA (GGC Gly	6855
Ile	GGT Gly 140	GTT Val	CAG Gln	CTT Leu	AAA (Lys 1 2:	SAC A Asp I 145	TC A le L	AG (GTC f	Phe l	AAC Asn 150	AGG Arg	GAC Asp	AGG Arg	CAG Gln	6903
AAG Lys 2155	GGT Gly	ACC Thr	AAT Asn	Val	TGC (Cys 1 160	GCG G Ala V	TA G	SCC A	Asn (GGC Gly 165	Gly	TGC Cys	CAG Gln	Gln	CTC Leu 170	6951
TGC Cys	TTG Leu	TAT Tyr	Arg	GGT Gly 2175	GGC (GGA C Gly G	AG (Arg .	GCC Ala 180	TGT Cys	GCC Ala	TGT Cys	Ala	CAC His 185	GJA GGG	6999
ATG Met	CTG Leu	Ala	GAA Glu 2190	GAC Asp	GGG Gly	GCC T Ala S	Ser (TGC Cys 195	CGA Arg	GAG Glu	TAC Tyr	Ala	GGC Gly 2200	TAC Tyr	CTG Leu	7047
CTC Leu	Tyr	TCA Ser 2205	GAG Glu	CGG Arg	ACC Thr	Ile I	CTC : Leu : 210	AAG Lys	AGC Ser	ATC Ile	His	CTG Leu 2215	TCG Ser	GAT Asp	GAG Glu	7095
Arg	AAC Asn 2220	CTC Leu	AAC Asn	GCA Ala	CCG Pro	GTG (Val (225	CAG Gln	CCC Pro	TTT Phe	Glu	GAC Asp 2230	CCC Pro	GAG Glu	CAC His	ATG Met	7143
AAA Lys 2235	Asn	GTC Val	ATC Ile	Ala	CTG Leu 2240	GCC '	TTT Phe	GAC Asp	Tyr	CGA Arg 2245	GCA Ala	GGC Gly	ACC Thr	Ser	CCG Pro 2250	7191
GGG Gly	ACC Thr	CCT Pro	Asn	CGC Arg 2255	ATC Ile	TTC Phe	TTC Phe	Ser	GAC Asp 2260	ATC Ile	CAC His	TTT Phe	GIÀ	AAC Asn 2265	TTE	7239
CAG Gln	CAG Gln	ATC	AAT Asn 2270	Asp	GAT Asp	GGC Gly	Ser	GGC Gly 2275	Arg	ACC Thr	ACC	lle	GTG Val 2280	GIU	AAT Asn	7287 •
GTG Val	GGC Gly	TCT Ser 2285	· Val	GAA Glu	GGC	Leu	GCC Ala 290	TAT Tyr	CAC His	CGT Arg	GG(TGG Trp 2295) Asp	ACA Thr	CTG Leu	7335
TAC Tyr	TGC Trp 2300	Thi	A AGO	TAC	Thr	ACA Thr 2305	TCC Ser	ACC Thr	: ATC	ACC Thr	CGC Arc 2310	g His	C ACC	C GTC	GAC Asp	7383
CA(Gl: 231:	n Thi	r CG	c cci g Pro	A GGC	GCC Ala 2320	Phe	GAG Glu	AGG	GAG Glu	ACF 1 Thi 2325	· Va	C ATO	C ACC	C ATO	TCC Ser 2330	7431
GG;	A GAG y Ası	C GA p As	C CA	C CCC s Pro 233	o Arg	GCC Ala	TTT	GT(G CT(Lev 2340	u Ası	r GA p Gl	G TG u Cy	C CA s Gl	G AAG n As: 234	C CTG n Leu 5	7479
AT Me	G TT	C TG e Tr	G AC p Th 235	r As	T TGC n Trp	AAC Asn	GAC Glu	CTC Let 235	u Hi	T CC.	A AG o Se	C AT	C AT e Me 236	t Ar	G GCA g Ala	··· 7527

(Sh et 31 of 91)

GCC (Leu					Val					Glu					7575
					Ala					Ala		AAG Lys				7623
				Leu					Arg			TAC Tyr		Gly		7671
			Val					Glu				CCC Pro	Phe			7719
-		Tyr					Phe					GTG Val				7767
	Gln					Tyr					Met	AAG Lys 2455				7815
Val					Gln					Ile		GTG Val				7863
				Glu					Arg			AAT Asn		Gly		7911
			Cys					Gln		His		AAC Asn	Cys			7959
		Gly					Glu		Phe			CGG Arg				8007
	Ser					Asp		Phe			Ala	AAT Asn 2535				8055
Ile					Thr		Asp			Ser		TGC Cys				8103
				Pro		Tyr					Arg	TGC Cys		Lys		8151
			Cys		Asn					Ser		ATG Met			Cys	8199
AAT Asn	G1 A	GTG Val	GAT Asp 2590	Tyr	TGT Cys	GGG Gly	GA7	GGC Gly 259	y Sex	TAD 1	GAC Glu	S ATA	CC1 Pro 2600	Cys	AAC Asn	8247

•	
AAG ACT GCC TGT GGT GTG GGT GAG TTC CGC TGC CGG GAT GGG TCC TGC Lys Thr Ala Cys Gly Val Gly Glu Phe Arg Cys Arg Asp Gly Ser Cys 2605 2610 2615	8295
ATC GGG AAC TCC AGT CGC TGC AAC CAG TTT GTG GAT TGT GAG GAT GCC Ile Gly Asn Ser Ser Arg Cys Asn Gln Phe Val Asp Cys Glu Asp Ala 2620 2625 2630	8343
TCG GAT GAG ATG AAT TGC AGT GCC ACA GAC TGC AGC AGC TAT TTC CGC Ser Asp Glu Met Asn Cys Ser Ala Thr Asp Cys Ser Ser Tyr Phe Arg 2635	8391
CTG GGC GTG AAA GGT GTC CTC TTC CAG CCG TGC GAG CGG ACA TCC CTG Leu Gly Val Lys Gly Val Leu Phe Gln Pro Cys Glu Arg Thr Ser Leu 2655 2660 2665	8439
TGC TAC GCA CCT AGC TGG GTG TGT GAT GGC GCC AAC GAC TGT GGA GAC Cys Tyr Ala Pro Ser Trp Val Cys Asp Gly Ala Asn Asp Cys Gly Asp 2670 2680	8487
TAC AGC GAT GAA CGT GAC TGT CCA GGT GTG AAG CGC CCT AGG TGC CCG Tyr Ser Asp Glu Arg Asp Cys Pro Gly Val Lys Arg Pro Arg Cys Pro 2695 2690 2695	8535
CTC AAT TAC TTT GCC TGC CCC AGC GGG CGC TGT ATC CCC ATG AGC TGG Leu Asn Tyr Phe Ala Cys Pro Ser Gly Arg Cys Ile Pro Met Ser Trp 2700 2705 2710	8583
ACG TGT GAC AAG GAG GAT GAC TGT GAG AAC GGC GAG GAT GAG ACC CAC Thr Cys Asp Lys Glu Asp Asp Cys Glu Asn Gly Glu Asp Glu Thr His 2715 2720 2725	8631
TGC AAC AAG TTC TGC TCA GAG GCA CAG TTC GAG TGC CAG AAC CAC CGG Cys Asn Lys Phe Cys Ser Glu Ala Gln Phe Glu Cys Gln Asn His Arg 2735 2740 2745	8679
TGT ATC TCC AAG CAG TGG CTG TGT GAC GGT AGC GAT GAT TGC GGG GAT Cys Ile Ser Lys Gln Trp Leu Cys Asp Gly Ser Asp Asp Cys Gly Asp 2750 2755 2760	8727
GGC TCC GAT GAG GCA GCT CAC TGT GAA GGC AAG ACA TGT GGC CCC TCC Gly Ser Asp Glu Ala Ala His Cys Glu Gly Lys Thr Cys Gly Pro Ser 2765 2770 2775	8775
TCC TTC TCC TGT CCC GGC ACC CAC GTG TGT GTC CCT GAG CGC TGG CTC Ser Phe Ser Cys Pro Gly Thr His Val Cys Val Pro Glu Arg Trp Let 2780 2780	8823
TGT GAT GGC GAC AAG GAC TGT ACC GAT GGC GCG GAT GAG AGT GTC ACC Cys Asp Gly Asp Lys Asp Cys Thr Asp Gly Ala Asp Glu Ser Val The 2795 2800 2805	•
GCT GGC TGC CTG TAC AAC AGC ACC TGT GAT GAC CGT GAG TTC ATG TG Ala Gly Cys Leu Tyr Asn Ser Thr Cys Asp Asp Arg Glu Phe Met Cy 2815 2820 2825	C 8919 s
CAG AAC CGC TTG TGT ATT CCC AAG CAT TTC GTG TGC GAC CAT GAC CG Gln Asn Arg Leu Cys Ile Pro Lys His Phe Val Cys Asp His Asp Ar 2830 2835 2840	T 8967

(Sh et 33 of 91)

GAC TGT GCT GAT GGC TCT GAT GAA TCC CCT GAG TGT GAG TAC CCA ACC Asp Cys Ala Asp Gly Ser Asp Glu Ser Pro Glu Cys Glu Tyr Pro Thr 2845 2850 2855	9015
TGC GGG CCC AAT GAA TTC CGC TGT GCC AAT GGG CGT TGT CTG AGC TCC Cys Gly Pro Asn Glu Phe Arg Cys Ala Asn Gly Arg Cys Leu Ser Ser 2860 2865 2870	9063
CGT CAG TGG GAA TGT GAT GGG GAG AAT GAC TGT CAC GAC CAC AGC GAT Arg Gln Trp Glu Cys Asp Gly Glu Asn Asp Cys His Asp His Ser Asp 2875 2880 2885	9111
GAG GCT CCC AAG AAC CCA CAC TGC ACC AGC CCA GAG CAC AAA TGC AAT Glu Ala Pro Lys Asn Pro His Cys Thr Ser Pro Glu His Lys Cys Asn 2900 2905	9159
GCC TCA TCA CAG TTC CTG TGC AGC AGC GGG CGC TGC GTG GCT GAG GCG Ala Ser Ser Gln Phe Leu Cys Ser Ser Gly Arg Cys Val Ala Glu Ala 2910 2915 2920	9207
TTG CTC TGC AAC GGC CAG GAC GAC TGT GGG GAC GGT TCA GAC GAA CGC Leu Leu Cys Asn Gly Gln Asp Asp Cys Gly Asp Gly Ser Asp Glu Arg 2925 2930	9255
GGG TGC CAT GTC AAC GAG TGT CTC AGC CGC AAG CTC AGT GGC TGC AGT Gly Cys His Val Asn Glu Cys Leu Ser Arg Lys Leu Ser Gly Cys Ser 2940 2950	9303
CAG GAC TGC GAG GAC CTC AAG ATA GGC TTT AAG TGC CGC TGT CGC CCG Gln Asp Cys Glu Asp Leu Lys Ile Gly Phe Lys Cys Arg Cys Arg Pro 2955 2960 2965	9351
GGC TTC CGG CTA AAG GAC GAT GGC AGG ACC TGT GCC GAC CTG GAT GAG Gly Phe Arg Leu Lys Asp Asp Gly Arg Thr Cys Ala Asp Leu Asp Glu 2975 2980 2985	9399
TGC AGC ACC ACC TTC CCC TGC AGC CAG CTC TGC ATC AAC ACC CAC GGA Cys Ser Thr Thr Phe Pro Cys Ser Gln Leu Cys Ile Asn Thr His Gly 2990 2995 3000	9447 .
AGT TAC AAG TGT CTG TGT GTG GAG GGC TAT GCA CCC CGT GGC GGT GAC Ser Tyr Lys Cys Leu Cys Val Glu Gly Tyr Ala Pro Arg Gly Gly Asp 3005 3010 3015	9495
CCC CAC AGC TGC AAA GCT GTG ACC GAT GAG GAG CCA TTT CTC ATC TTT Pro His Ser Cys Lys Ala Val Thr Asp Glu Glu Pro Phe Leu Ile Phe 3020 3025 3030	9543
GCC AAC CGG TAC TAC CTG CGG AAG CTC AAC CTG GAC GGC TCC AAC TAC Ala Asn Arg Tyr Tyr Leu Arg Lys Leu Asn Leu Asp Gly Ser Asn Tyr 3035 3040 3045	9591
ACA CTG CTT AAG CAG GGC CTG AAC AAT GCG GTC GCC TTG GCA TTT GAC Thr Leu Leu Lys Gln Gly Leu Asn Asn Ala Val Ala Leu Ala Phe Asp 3055 3060 3065	.9639
TAC CGA GAG CAG ATG ATC TAC TGG ACG GGC GTG ACC ACC CAG GGC AGC Tyr Arg Glu Gln Met Ile Tyr Trp Thr Gly Val Thr Thr Gln Gly Ser 3070 3080	9687

(Sh et 34 of 91)

ATG ATT CGC AGG ATG CAC CTC AAC GGC AGC AAC GTG CAG GTT CTG CAC Met Ile Arg Arg Met His Leu Asn Gly Ser Asn Val Gln Val Leu His 3085	9735
CGG ACG GGC CTT AGT AAC CCA GAT GGG CTC GCT GTG GAC TGG GTG GGT Arg Thr Gly Leu Ser Asn Pro Asp Gly Leu Ala Val Asp Trp Val Gly 3100 3105 3110	9783
GGC AAC CTG TAC TGG TGT GAC AAG GGC AGA GAT ACC ATT GAG GTG TCC Gly Asn Leu Tyr Trp Cys Asp Lys Gly Arg Asp Thr Ile Glu Val Ser 3115 3120 3125 3130	9831
AAG CTT AAC GGG GCC TAT CGG ACA GTG CTG GTC AGC TCT GGC CTC CGG Lys Leu Asn Gly Ala Tyr Arg Thr Val Leu Val Ser Ser Gly Leu Arg 3135 3140 3145	9879
GAG CCC AGA GCT CTG GTA GTG GAT GTA CAG AAT GGG TAC CTG TAC TGG Glu Pro Arg Ala Leu Val Val Asp Val Gln Asn Gly Tyr Leu Tyr Trp 3150 3155 3160	9927
ACA GAC TGG GGT GAC CAC TCA CTG ATC GGC CGG ATT GGC ATG GAT GGA Thr Asp Trp Gly Asp His Ser Leu Ile Gly Arg Ile Gly Met Asp Gly 3165 3170 3175	9975
TCT GGC CGC AGC ATC ATC GTG GAC ACT AAG ATC ACA TGG CCC AAT GGC Ser Gly Arg Ser Ile Ile Val Asp Thr Lys Ile Thr Trp Pro Asn Gly 3180 3185 3190	10023
CTG ACC GTG GAC TAC GTC ACG GAA CGC ATC TAC TGG GCT GAC GCC CGT Leu Thr Val Asp Tyr Val Thr Glu Arg Ile Tyr Trp Ala Asp Ala Arg 3195 3200 3205 3210	10071
GAG GAC TAC ATC GAG TTC GCC AGC CTG GAT GGC TCC AAC CGT CAC GTT Glu Asp Tyr Ile Glu Phe Ala Ser Leu Asp Gly Ser Asn Arg His Val 3215 3220 3225	10119
GTG CTG AGC CAA GAC ATC CCA CAC ATC TTT GCG CTG ACC CTA TTT GAA Val Leu Ser Gln Asp Ile Pro His Ile Phe Ala Leu Thr Leu Phe Glu 3230 3235 3240	10167 .
GAC TAC GTC TAC TGG ACA GAC TGG GAA ACG AAG TCC ATC AAC CGG GCC Asp Tyr Val Tyr Trp Thr Asp Trp Glu Thr Lys Ser Ile Asn Arg Ala 3245 3250 3255	10215
CAC AAG ACC ACG GGT GCC AAC AAA ACA CTC CTC ATC AGC ACC CTG CAC His Lys Thr Thr Gly Ala Asn Lys Thr Leu Leu Ile Ser Thr Leu His 3260 3265 3270	10263
CGG CCC ATG GAC TTA CAT GTA TTC CAC GCC CTG CGC CAG CCA GAT GTG Arg Pro Met Asp Leu His Val Phe His Ala Leu Arg Gln Pro Asp Val 3275 3280 3285 3290	10311
CCC AAT CAC CCC TGC AAA GTC AAC AAT GGT GGC TGC AGC AAC CTG TGC Pro Asn His Pro Cys Lys Val Asn Asn Gly Gly Cys Ser Asn Leu Cys 3295 3300 3305	10359
CTG CTG TCC CCT GGG GGT GGT CAC AAG TGC GCC TGC CCC ACC AAC TTC Leu Leu Ser Pro Gly Gly His Lys Cys Ala Cys Pro Thr Asn Phe 3310 3320	10407

FIG. 12A

(Sheet 35 of 91)

8449-134		(Sheet 35 of 91)
	GC CGT ACC TGT GTG TCC I Ly Arg Thr Cys Val Ser I 3330	
-	AT GAC AAG TGC ATC CCC on Asp Lys Cys Ile Pro 3	
GAC ACG GAG GAC GAC TO	GT GGG GAT CAC TCA GAC	GAG CCT CCA GAC TGT 10551
Asp Thr Glu Asp Asp Cy	ys Gly Asp His Ser Asp	Glu Pro Pro Asp Cys
3355	60 3365	3370
CCC GAG TTC AAG TGC CC	GC CCA GGC CAG TTC CAG	TGC TCC ACC GGC ATC 10599
Pro Glu Phe Lys Cys Au	rg Pro Gly Gln Phe Gln	Cys Ser Thr Gly Ile
3375	3380	3385
	TC ATC TGT GAT GGG GAC he Ile Cys Asp Gly Asp 3395	
AAT AGT GAC GAG GCC AN	AT TGC GAC ATT CAC GTC	TGC TTG CCC AGC CAA 10695
Asn Ser Asp Glu Ala As	sn Cys Asp Ile His Val	Cys Leu Pro Ser Gln
3405	3410	3415
TTC AAG TGC ACC AAC AG	CC AAC CGC TGC ATT CCT	GGC ATC TTC CGT TGC 10743
Phe Lys Cys Thr Asn T	hr Asn Arg Cys Ile Pro	Gly Ile Phe Arg Cys
3420	3425	3430
AAT GGG CAG GAC AAC TO	GC GGG GAC GGC GAG GAT	GAG CGG GAT TGC CCT 10791
Asn Gly Gln Asp Asn C	ys Gly Asp Gly Glu Asp	Glu Arg Asp Cys Pro
3435	40 3445	3450
GAG GTG ACC TGC GCC C	CC AAC CAG TTC CAG TGC	TCC ATC ACC AAG CGC 10839
Glu Val Thr Cys Ala P	TO Asn Gln Phe Gln Cys	Ser Ile Thr Lys Arg
3455	3460	3465
TGC ATC CCT CGC GTC T	rgg GTC TGT GAC AGG GAT	AAT CAC TGT GTG GAC 10887
Cys Ile Pro Arg Val T	rp Val Cys Asp Arg Asp	Asn His Cys Val Asp
3470	3475	3480
GGC AGT GAT GAG CCT G	SCC AAC TGT ACC CAA ATG	ACC TGT GGA GTG GAT 10935
Gly Ser Asp Glu Pro A	Ala Asn Cys Thr Gln Met	Thr Cys Gly Val Asp
3485	3490	3495
GAG TTC CGC TGC AAG G	GAT TCT GGC CGC TGC ATC	CCC GCG CGC TGG AAG 10983
Glu Phe Arg Cys Lys A	Asp Ser Gly Arg Cys Ile	Pro Ala Arg Trp Lys
3500	3505	3510
Cys Asp Gly Glu Asp A	GAC TGT GGG GAT GGT TCA Asp Cys Gly Asp Gly Ser 520 3525	Asp Glu Pro Lys Glu
GAG TGT GAT GAG CGC A Glu Cys Asp Glu Arg 3 3535	ACC TGT GAG CCA TAC CAG Thr Cys Glu Pro Tyr Gln 3540	TTC CGC TGC AAA AAC 11079 Phe Arg Cys Lys Asn 3545
AAC CGC TGT GTC CCA	GGC CGT TGG CAA TGT GAC	TAC GAC AAC GAC TGC 11127

FIG. 12A

3560

Asn Arg Cys Val Pro Gly Arg Trp Gln Cys Asp Tyr Asp Asn Asp Cys

3550 ·

3555

8449-134 (Sheet 36 of 91)

	GAT Asp 3					Glu					Arg					11175
Ser	GAG Glu 3580	Phe			Ala					Ile						11223
	GAT Asp			His					Gly					Asp		11271
	CCC Pro		Cys					Phe					Gly		TGC Cys	11319
	CCC Pro	Leu					Asp					Cys				11367
	GAC Asp					Gly					Thr					11415
Glu	TTT Phe 3660				Asn					Pro						11463
	Gly			Asp					Ser		Glu			Glu		11511
	GCC Ala		Phe					Asn					Cys		AAT Asn	11559
	CGA Arg	Val					Gly					Gly		Asp		11607 .
TGT Cys	Gly	GAT Asp 3725	GGG Gly	ACT Thr	GAC Asp	Glu	GAG Glu 3730	Asp	TGT Cys	GAG Glu	Pro	CCC Pro 3735	Thr	GCC Ala	CAG Gln	11655
Asn	CCC Pro 3740	His			Asp		Lys			Leu		Arg			CGC	11703
	Leu			Ser		Arg			Met		Asp				GAT Asp 3770	11751 · .
					Asp					Pro					Cys	11799
GCC Ala	ACC Thr	AAT Asn	GCC Ala 3790	Ser	ATG Met	TG1 Cys	: Gl7	GAC Asp 3795	o Glu	GCI Ala	CGT Aro	TG1 G Cys	GT(S Va: 380	l Ar	C ACT	11847

(She t 37 of 91) 8449-134 11895 GAG AAA GCT GCC TAC TGT GCC TGC CGC TCG GGC TTC CAT ACT GTG CCG Glu Lys Ala Ala Tyr Cys Ala Cys Arg Ser Gly Phe His Thr Val Pro 3810 3815 3805 GGC CAG CCC GGA TGC CAG GAC ATC AAC GAG TGC CTG CGC TTT GGT ACC 11943 Gly Gln Pro Gly Cys Gln Asp Ile Asn Glu Cys Leu Arg Phe Gly Thr 3825 TGC TCT CAG CTC TGG AAC AAA CCC AAG GGA GGC CAC CTC TGC AGC TGT 11991 -Cys Ser Gln Leu Trp Asn Lys Pro Lys Gly Gly His Leu Cys Ser Cys 3840 3845 3835 GCC CGC AAC TTC ATG AAG ACA CAC AAC ACC TGC AAA GCT GAA GGC TCC 12039 Ala Arg Asn Phe Met Lys Thr His Asn Thr Cys Lys Ala Glu Gly Ser 3865 3855 12087 GAG TAC CAG GTG CTA TAC ATC GCG GAT GAC AAC GAG ATC CGC AGC TTG Glu Tyr Gln Val Leu Tyr Ile Ala Asp Asp Asn Glu Ile Arg Ser Leu 3875 3870 TTC CCG GGC CAC CCC CAC TCA GCC TAC GAG CAG ACA TTC CAG GGC GAT 12135 Phe Pro Gly His Pro His Ser Ala Tyr Glu Gln Thr Phe Gln Gly Asp 3895 3890 3885 GAG AGT GTC CGC ATA GAT GCC ATG GAT GTC CAT GTC AAG GCC GGC CGT 12183 Glu Ser Val Arg Ile Asp Ala Met Asp Val His Val Lys Ala Gly Arg 3905 3910 3900 GTC TAC TGG ACT AAC TGG CAC ACG GGC ACA ATC TCC TAC AGG AGC CTG 12231 Val Tyr Trp Thr Asn Trp His Thr Gly Thr Ile Ser Tyr Arg Ser Leu 3925 3920 3915_. CCC CCT GCC GCC CCT CCT ACC ACT TCC AAC CGC CAC CGG AGG CAG ATC 12279 Pro Pro Ala Ala Pro Pro Thr Thr Ser Asn Arg His Arg Arg Gln Ile 3945 3940 3935 GAC CGG GGT GTC ACC CAC CTC AAT ATT TCA GGG CTG AAG ATG CCG AGG 12327 . Asp Arg Gly Val Thr His Leu Asn Ile Ser Gly Leu Lys Met Pro Arg 3960 3955 3950 GGT ATC GCT ATC GAC TGG GTG GCC GGG AAT GTG TAC TGG ACC GAT TCC 12375 Gly Ile Ala Ile Asp Trp Val Ala Gly Asn Val Tyr Trp Thr Asp Ser

ACG CTC ATC TCG GGC ATG ATT GAT GAG CCC CAT GCC ATC GTG GTG GAC Thr Leu Ile Ser Gly Met Ile Asp Glu Pro His Ala Ile Val Val Asp 4000 4005 3995 CCT CTG AGG GGC ACC ATG TAC TGG TCA GAC TGG GGG AAC CAC CCC AAG 12519 Pro Leu Arg Gly Thr Met Tyr Trp Ser Asp Trp Gly Asn His Pro Lys 4025 4020 4015 ATT GAA ACA GCA GCG ATG GAT GGC ACC CTT CGG GAG ACT CTC GTG CAA 12567 Ile Glu Thr Ala Ala Met Asp Gly Thr Leu Arg Glu Thr Leu Val Gln 4040 4035 4030

3970

3985

GGC CGA GAC GTG ATT GAG GTG GCG CAA ATG AAG GGC GAG AAC CGC AAG

Gly Arg Asp Val Ile Glu Val Ala Gln Met Lys Gly Glu Asn Arg Lys

3965

3980

3975

12423

12471

(Sheet 38 of 91)

8449-134		(Sheet 38 of 9
GAC AAC ATT CAG TGG CO Asp Asn Ile Gln Trp Po 4045	CT ACA GGG CTG GCT GTG ro Thr Gly Leu Ala Val 4050	GAC TAT CAC AAT GAA 12615 Asp Tyr His Asn Glu 4055
CGG CTC TAC TGG GCA GA Arg Leu Tyr Trp Ala A: 4060	AT GCC AAG CTT TCG GTC sp Ala Lys Leu Ser Val	ATC GGC AGC ATC CGG 12663 Ile Gly Ser Ile Arg 070
CTC AAC GGC ACT GAC CO Leu Asn Gly Thr Asp Pt 4075	CC ATT GTG GCT GCT GAC ro Ile Val Ala Ala Asp 4085	AGC AAA CGA GGC CTA 12711 Ser Lys Arg Gly Leu 4090
AGT CAC CCC TTC AGC AGS Ser His Pro Phe Ser I. 4095	TC GAT GTG TTT GAA GAC le Asp Val Phe Glu Asp 4100	TAC ATC TAC GGA GTC 12759 Tyr Ile Tyr Gly Val 4105
ACT TAC ATC AAT AAT CO Thr Tyr Ile Asn Asn As 4110	GT GTC TTC AAG ATC CAC rg Val Phe Lys Ile His 4115	AAG TTT GGA CAC AGC 12807 Lys Phe Gly His Ser 4120
CCC TTG TAC AAC CTA AG Pro Leu Tyr Asn Leu Tl 4125	CT GGG GGC CTG AGC CAT hr Gly Gly Leu Ser His 4130	GCC TCT GAT GTA GTC 12855 Ala Ser Asp Val Val 4135
CTT TAC CAT CAA CAC AN Leu Tyr His Gln His Ly 4140	AG CAG CCT GAA GTG ACC ys Gln Pro Glu Val Thr 4145	AAC CCC TGT GAC CGC 12903 Asn Pro Cys Asp Arg
	TG TGT CTG CTG AGC CCC eu Cys Leu Leu Ser Pro 60 4165	
ACC TGT CCC AAT GGA AND Thr Cys Pro Asn Gly Ly 4175	AG AGG CTG GAT AAT GGC ys Arg Leu Asp Asn Gly 4180	ACC TGT GTG CCT GTG 12999 Thr Cys Val Pro Val 4185
CCC TCT CCA ACA CCC CC Pro Ser Pro Thr Pro P. 4190	CT CCA GAT GCC CCT AGG ro Pro Asp Ala Pro Arg 4195	CCT GGA ACC TGC ACT 13047 Pro Gly Thr Cys Thr 4200
	GT GGT AGT TGT TTC CTC ly Gly Ser Cys Phe Leu 4210	
	AG CCC CGT TAC ACA GGC ln Pro Arg Tyr Thr Gly 4225	
	AC TGT CAC AAC GGA GGC yr Cys His Asn Gly Gly 40 4245	
CCA TCT GGC ATG CCC A Pro Ser Gly Met Pro T 4255	CG TGC CGC TGT CCC ACT hr Cys Arg Cys Pro Thr 4260	GGC TTC ACG GGC CCC 13239 Gly Phe Thr Gly Pro 4265
	TG TGT GCA GGC TAC TGC al Cys Ala Gly Tyr Cys 4275	

FIG. 12A

4280

4275

(Sheet 39 of 91)

8449-134

TGC ACC GTC AAC CAG GGC AAC CAG CCC CAG TGC CGA TGT CTA CCT GGC Cys Thr Val Asn Gln Gly Asn Gln Pro Gln Cys Arg Cys Leu Pro Gly 4285	13335
TTC CTG GGC GAC CGT TGC CAG TAC CGG CAG TGC TCT GGC TTC TGT GAG Phe Leu Gly Asp Arg Cys Gln Tyr Arg Gln Cys Ser Gly Phe Cys Glu 4300 4305 4310	13383
AAC TTT GGC ACC TGT CAG ATG GCT GCT GAT GGC TCC CGA CAA TGT CGC Asn Phe Gly Thr Cys Gln Met Ala Ala Asp Gly Ser Arg Gln Cys Arg 4325	13431
TGC ACC GTC TAC TTT GAG GGA CCA AGG TGT GAG GTG AAC AAG TGT AGT Cys Thr Val Tyr Phe Glu Gly Pro Arg Cys Glu Val Asn Lys Cys Ser 4335 4340 4345	13479
CGC TGT CTC CAA GGC GCC TGT GTG GTC AAT AAG CAG ACC GGA GAT GTC Arg Cys Leu Gln Gly Ala Cys Val Val Asn Lys Gln Thr Gly Asp Val 4350 4360	13527
ACA TGC AAC TGC ACT GAT GGC CGG GTA GCC CCC AGT TGT CTC ACC TGC Thr Cys Asn Cys Thr Asp Gly Arg Val Ala Pro Ser Cys Leu Thr Cys 4365 4370 4375	13575
ATC GAT CAC TGT AGC AAT GGT GGC TCC TGC ACC ATG AAC AGC AAG ATG Ile Asp His Cys Ser Asn Gly Gly Ser Cys Thr Met Asn Ser Lys Met 4380 4385 4390	13623
ATG CCT GAG TGC CAG TGC CCG CCC CAT ATG ACA GGA CCC CGG TGC CAG Met Pro Glu Cys Gln Cys Pro Pro His Met Thr Gly Pro Arg Cys Gln 4395 4400 4410	13671
GAG CAG GTT GTT AGT CAG CAA CAG CCT GGG CAT ATG GCC TCC ATC CTG Glu Gln Val Val Ser Gln Gln Gln Pro Gly His Met Ala Ser Ile Leu 4415 4420 4425	13719
ATC CCT CTG CTG CTT CTC CTG CTG CTT CTG GTG G	13767 .
TTC TGG TAT AAG CGG CGA GTC CGA GGG GCT AAG GGC TTC CAG CAC CAG Phe Trp Tyr Lys Arg Arg Val Arg Gly Ala Lys Gly Phe Gln His Gln 4455 4455	13815
CGG ATG ACC AAT GGG GCC ATG AAT GTG GAA ATT GGA AAC CCT ACC TAC Arg Met Thr Asn Gly Ala Met Asn Val Glu Ile Gly Asn Pro Thr Tyr 4460 4465 4470	13863
AAG ATG TAT GAA GGT GGA GAG CCC GAT GAT GTC GGG GGC CTA CTG GAT Lys Met Tyr Glu Gly Glu Pro Asp Asp Val Gly Gly Leu Leu Asp 4475 4480 4485	13911
GCT GAT TTT GCC CTT GAC CCT GAC AAG CCT ACC AAC TTC ACC AAC CCA Ala Asp Phe Ala Leu Asp Pro Asp Lys Pro Thr Asn Phe Thr Asn Pro 4505	
GTG TAT GCC ACG CTC TAC ATG GGG GGC CAC GGC AGC CGC CAT TCC CTG Val Tyr Ala Thr Leu Tyr Met Gly Gly His Gly Ser Arg His Ser Leu 4510 4520	14007

14650

14710

14770

14830

GCC AGC ACG GAC GAG AAG CGA GAA CTG CTG GGC CGG GGA CCT GAA GAC 14055 Ala Ser Thr Asp Glu Lys Arg Glu Leu Leu Gly Arg Gly Pro Glu Asp 4525 GAG ATA GGA GAT CCC TTG GCA TAGGGCCCTG CCCCGACGGA TGTCCCCAGA AAGC 14110 CCCCTGCCAC ATGAGTCTTT CAATGAACCC CCTCCCCAGC CGGCCCTTCT CCGGCCCTGC Glu Ile Gly Asp Pro Leu Ala 4540 CGGGTGTACA AATGTAAAAA TGAAGGAATT ACTTTTTATA TGTGAGCGAG CAAGCGAGCA 14230 AGCACAGTAT TATCTCTTTG CATTTCCTTC CTGCCTGCTC CTCAGTATCC CCCCCATGCT GCCTTGAGGG GGCGGGGAGG GCTTTGTGGC TCAAAGGTAT GAAGGAGTCC ACATGTTCCC 14350 TACCGAGCAT ACCCCTGGAA GCCTGGCGGC ACGGCCTCCC CACCACGCCT GTGCAAGACA 14410 CTCAACGGGG CTCCGTGTCC CAGCTTTCCT TTCCTTGGCT CTCTGGGGTT AGTTCAGGGG 14470 AGGTGGAGTC CTCTGCTGAC CCTGTCTGGA AGATTTGGCT CTAGCTGAGG AAGGAGTCTT 14530 TTAGTTGAGG GAAGTCACCC CAAACCCCAG CTCCCACTTT CAGGGGCACC TCTCAGATGG 14590

CCATGCTCAG TATCCCTTCC AGACAGGCCC TCCCCTCTCT AGCGCCCCCT CTGTGGCTCC

TAGGGCTGAA CACATTCTTT GGTAACTGTC CCCCAAGCCT CCCATCCCCC TGAGGGCCAG

GAAGAGTCGG GGCACACCAA GGAAGGGCAA GCGGGCAGCC CCATTTTGGG GACGTGAACG

TTTTAATAAT TTTTGCTGAA TTCCTTTACA ACTAAATAAC ACAGATATTG TTATAAATAA AATTGTAAAA AAAAAAAAAA

Met Leu Thr Pro Pro Leu Leu Leu Val Pro Leu Leu Ser Ala Leu Val Ser Gly Ala Thr Met Asp Ala Pro Lys Thr Cys Ser Pro Lys Gln Phe Ala Cys Arg Asp Gln Ile Thr Cys Ile Ser Lys Gly Trp Arg Cys Asp Gly Glu Arg Asp Cys Pro Asp Gly Ser Asp Glu Ala Pro Glu Ile Cys Pro Gln Ser Lys Ala Gln Arg Cys Pro Pro Asn Glu His Ser Cys 75 Leu Gly Thr Glu Leu Cys Val Pro Met Ser Arg Leu Cys Asn Gly Ile Gln Asp Cys Met Asp Gly Ser Asp Glu Gly Ala His Cys Arg Glu Leu Arg Ala Asn Cys Ser Arg Met Gly Cys Gln His His Cys Val Pro Thr 120 Pro Ser Gly Pro Thr Cys Tyr Cys Asn Ser Ser Phe Gln Leu Glu Ala 135 Asp Gly Lys Thr Cys Lys Asp Phe Asp Glu Cys Ser Val Tyr Gly Thr 150 155 Cys Ser Gln Leu Cys Thr Asn Thr Asp Gly Ser Phe Thr Cys Gly Cys Val Glu Gly Tyr Leu Leu Gln Pro Asp Asn Arg Ser Cys Lys Ala Lys 185 Asn Glu Pro Val Asp Arg Pro Pro Val Leu Leu Ile Ala Asn Ser Gln Asn Ile Leu Ala Thr Tyr Leu Ser Gly Ala Gln Val Ser Thr Ile Thr 215 Pro Thr Ser Thr Arg Gln Thr Thr Ala Met Asp Phe Ser Tyr Ala Asn 230 Glu Thr Val Cys Trp Val His Val Gly Asp Ser Ala Ala Gln Thr Gln 245 250 Leu Lys Cys Ala Arg Met Pro Gly Leu Lys Gly Phe Val Asp Glu His Thr Ile Asn Ile Ser Leu Ser Leu His His Val Glu Gln Met Ala Ile 280 Asp Trp Leu Thr Gly Asn Phe Tyr Phe Val Asp Asp Ile Asp Asp Arg 295 Ile Phe Val Cys Asn Arg Asn Gly Asp Thr Cys Val Thr Leu Leu Asp 310 315 Leu Glu Leu Tyr Asn Pro Lys Gly Ile Ala Leu Asp Pro Ala Met Gly 325 330 Lys Val Phe Phe Thr Asp Tyr Gly Gln Ile Pro Lys Val Glu Arg Cys Asp Met Asp Gly Gln Asn Arg Thr Lys Leu Val Asp Ser Lys Ile Val 360 365 355 Phe Pro His Gly Ile Thr Leu Asp Leu Val Ser Arg Leu Val Tyr Trp Ala Asp Ala Tyr Leu Asp Tyr Ile Glu Val Val Asp Tyr Glu Gly Lys 390 395 Gly Arg Gln Thr Ile Ile Gln Gly Ile Leu Ile Glu His Leu Tyr Gly 405 410 Leu Thr Val Phe Glu Asn Tyr Leu Tyr Ala Thr Asn Ser Asp Asn Ala 425 420 Asn Thr Gln Gln Lys Thr Ser Val Ile Arg Val Asn Arg Phe Asn Ser 440 Thr Glu Tyr Gln Val Val Thr Arg Val Asp Lys Gly Gly Ala Leu His 455 460 450

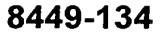
FIG. 12B

(Sheet 42 of 91)

0 -1			. •	•													•				
Ile	er	_ 1	ri e	Gl n	Aro	Ar	a (Gln	Pro	A c	ra	Val	Arc	Se	r	lis	Ala	Cy	<i>t</i> s (lu	
													- 4 -	•							
465 Asn						Ly	'S					470							-		
Ala				$\epsilon \alpha \alpha$	Lys	A)				_	רנונ	Arg	Cys				,	_			
Leu				Asp	Gl				~ /	() .		•				J & J					
Leu		1 '							Pr	0 (Зlу	Ile									
Gly	53	0	T	V=1	Pro	o A:	SD	Glu	Hi	s t	Met	Ilε	Pr	o I	le	G1u	As	n L	eu	Met	•
Asn						•							,					_			
Ala				E 0/	A						-) M -1							_			
				Th	r Il				n	111						~~~	,				
	_	_	Asp	Tr	р Ме																
	L				e Se			Ala	ı Aı												
	T				e G1	u G	ly														
					n G]	Ly 1						Th	r A								
					g A						Gli	ı Ar									
. 112	~ N	~~	675) 5 Tl	e P	ne V	/al	Th	r S	er	Ly	s Th	ır V	al I	Leu	Tr	p P	ro i	Asn	G)	. y
Le	u S	er	Lev	u As	p I	le !	Pro	Al	a G	ly	Ar	g Le	eu T 7	yr. ' 15	Trp	va	TH	Sp	WIO	72	20
70	5		_		Le G		710	, - Tl	_ T	.e.i	T.e	11 A	sn G	iv	Thr	As	рА	rg	Lys	I	Le
					/ Ly P 40																
					eu P																
					ly V																
	er (Glι	Ar, e		ro E																
					ly 7																
					eu /	Ala															
				al I	eu i																
			1 P		ro					Gl	n P										
<u>α</u>	, Sn	85 Ar	a C	vs]	(le	Gln	G)	u A	rg	Tr	p L	ys (Cys	Asp	G)	y A	.sp	Asn	As	p	Zys ถถก
					Ser																
					Arg 900																
T	rp	Le	eu C	ys . 15	900 Asp	Gly	A	sp A	Asn	As 92	9 Q	:ys	GLY	Asr	1 S	er (925	M9)	, ,		

Asn Ala Thr Cys Ser Ala Arg Thr Cys Pro Pro Asn Gln Phe Ser Cys Ala Ser Gly Arg Cys Ile Pro Ile Ser Trp Thr Cys Asp Leu Asp Asp Asp Cys Gly Asp Arg Ser Asp Glu Ser Ala Ser Cys Ala Tyr Pro Thr Cys Phe Pro Leu Thr Gln Phe Thr Cys Asn Asn Gly Arg Cys Ile Asn Ile Asn Trp Arg Cys Asp Asn Asp Asn Asp Cys Gly Asp Asn Ser Asp Glu Ala Gly Cys Ser His Ser Cys Ser Ser Thr Gln Phe Lys Cys Asn Ser Gly Arg Cys Ile Pro Glu His Trp Thr Cys Asp Gly Asp Asn Asp Cys Gly Asp Tyr Ser Asp Glu Thr His Ala Asn Cys Thr Asn Gln Ala Thr Arg Pro Pro Gly Gly Cys His Ser Asp Glu Phe Gln Cys Pro Leu Asp Gly Leu Cys Ile Pro Leu Arg Trp Arg Cys Asp Gly Asp Thr Asp Cys Met Asp Ser Ser Asp Glu Lys Ser Cys Glu Gly Val Thr His Val Cys Asp Pro Asn Val Lys Phe Gly Cys Lys Asp Ser Ala Arg Cys Ile Ser Lys Ala Trp Val Cys Asp Gly Asp Ser Asp Cys Glu Asp Asn Ser Asp Glu Glu Asn Cys Glu Ala Leu Ala Cys Arg Pro Pro Ser His Pro Cys Ala Asn Asn Thr Ser Val Cys Leu Pro Pro Asp Lys Leu Cys Asp . Gly Lys Asp Asp Cys Gly Asp Gly Ser Asp Glu Gly Glu Leu Cys Asp Gln Cys Ser Leu Asn Asn Gly Gly Cys Ser His Asn Cys Ser Val Ala Pro Gly Glu Gly Ile Val Cys Ser Cys Pro Leu Gly Met Glu Leu Gly Ser Asp Asn His Thr Cys Gln Ile Gln Ser Tyr Cys Ala Lys His Leu Lys Cys Ser Gln Lys Cys Asp Gln Asn Lys Phe Ser Val Lys Cys Ser Cys Tyr Glu Gly Trp Val Leu Glu Pro Asp Gly Glu Thr Cys Arg Ser Leu Asp Pro Phe Lys Leu Phe Ile Ile Phe Ser Asn Arg His Glu Ile Arg Arg Ile Asp Leu His Lys Gly Asp Tyr Ser Val Leu Val Pro Gly Leu Arg Asn Thr Ile Ala Leu Asp Phe His Leu Ser Gln Ser Ala Leu Tyr Trp Thr Asp Ala Val Glu Asp Lys Ile Tyr Arg Gly Lys Leu Leu Asp Asn Gly Ala Leu Thr Ser Phe Glu Val Val Ile Gln Tyr Gly Leu Ala Thr Pro Glu Gly Leu Ala Val Asp Trp Ile Ala Gly Asn Ile Tyr Trp Val Glu Ser Asn Leu Asp Gln Ile Glu Val Ala Lys Leu Asp Gly Thr Leu Arg Thr Thr Leu Leu Ala Gly Asp Ile Glu His Pro Arg Ala ... Ile Ala Leu Asp Pro Arg Asp Gly Ile Leu Phe Trp Thr Asp Trp Asp

Ala Ser Leu Pro Arg Ile Glu Ala Ala Ser Met Ser Gly Ala Gly Arg Arg Thr Ile His Arg Glu Thr Gly Ser Gly Gly Cys Ala Asn Gly Leu Thr Val Asp Tyr Leu Glu Lys Arg Ile Leu Trp Ile Asp Ala Arg Ser Asp Ala Ile Tyr Ser Ala Arg Tyr Asp Gly Ser Gly His Met Glu Val Leu Arg Gly His Glu Phe Leu Ser His Pro Phe Ala Val Thr Leu Tyr Gly Glu Val Tyr Trp Thr Asp Trp Arg Thr Asn Thr Leu Ala Lys Ala Asn Lys Trp Thr Gly His Asn Val Thr Val Val Gln Arg Thr Asn Thr Gln Pro Phe Asp Leu Gln Val Tyr His Pro Ser Arg Gln Pro Met Ala Pro Asn Pro Cys Glu Ala Asn Gly Gly Arg Gly Pro Cys Ser His Leu Cys Leu Ile Asn Tyr Asn Arg Thr Val Ser Trp Ala Cys Pro His Leu Met Lys Leu His Lys Asp Asn Thr Thr Cys Tyr Glu Phe Lys Lys Phe Leu Leu Tyr Ala Arg Gln Met Glu Ile Arg Gly Val Asp Leu Asp Ala Pro Tyr Tyr Asn Tyr Ile Ile Ser Phe Thr Val Pro Asp Ile Asp Asn Val Thr Val Leu Asp Tyr Asp Ala Arg Glu Gln Arg Val Tyr Trp Ser Asp Val Arg Thr Gln Ala Ile Lys Arg Ala Phe Ile Asn Gly Thr Gly Val Glu Thr Val Val Ser Ala Asp Leu Pro Asn Ala His Gly Leu Ala Val Asp Trp Val Ser Arg Asn Leu Phe Trp Thr Ser Tyr Asp Thr Asn Lys Lys Gln Ile Asn Val Ala Arg Leu Asp Gly Ser Phe Lys Asn Ala Val Val Gln Gly Leu Glu Gln Pro His Gly Leu Val His Pro Leu Arg Gly Lys Leu Tyr Trp Thr Asp Gly Asp Asn Ile Ser Met Ala Asn Met Asp Gly Ser Asn His Thr Leu Leu Phe Ser Gly Gln Lys Gly Pro Val Gly Leu Ala Ile Asp Phe Pro Glu Ser Lys Leu Tyr Trp Ile Ser Ser Gly Asn His Thr Ile Asn Arg Cys Asn Leu Asp Gly Ser Glu Leu Glu Val Ile Asp Thr Met Arg Ser Gln Leu Gly Lys Ala Thr Ala Leu Ala Ile Met Gly Asp Lys Leu Trp Trp Ala Asp Gln Val Ser Glu Lys Met Gly Thr Cys Asn Lys Ala Asp Gly Ser Gly Ser Val Val Leu Arg Asn Ser Thr Thr Leu Val Met His Met Lys Val Tyr Asp Glu Ser Ile Gln Leu Glu His Glu Gly Thr Asn Pro Cys Ser Val Asn Asn Gly Asp Cys Ser Gln Leu Cys Leu Pro Thr Ser Glu Thr Thr Arg Ser Cys



(Sheet 45 of 91)

Met Cys Thr Ala Gly Tyr Ser Leu Arg Ser Gly Gln Gln Ala Cys Glu Gly Val Gly Ser Phe Leu Leu Tyr Ser Val His Glu Gly Ile Arg Gly Ile Pro Leu Asp Pro Asn Asp Lys Ser Asp Ala Leu Val Pro Val Ser Gly Thr Ser Leu Ala Val Gly Ile Asp Phe His Ala Glu Asn Asp Thr Ile Tyr Trp Val Asp Met Gly Leu Ser Thr Ile Ser Arg Ala Lys Arg Asp Gln Thr Trp Arg Glu Asp Val Val Thr Asn Gly Ile Gly Arg Val Glu Gly Ile Ala Val Asp Trp Ile Ala Gly Asn Ile Tyr Trp Thr Asp Gln Gly Phe Asp Val Ile Glu Val Ala Arg Leu Asn Gly Ser Phe Arg Tyr Val Val Ile Ser Gln Gly Leu Asp Lys Pro Arg Ala Ile Thr Val His Pro Glu Lys Gly Tyr Leu Phe Trp Thr Glu Trp Gly His Tyr Pro Arg Ile Glu Arg Ser Arg Leu Asp Gly Thr Glu Arg Val Val Leu Val Asn Val Ser Ile Ser Trp Pro Asn Gly Ile Ser Val Asp Tyr Gln Gly Gly Lys Leu Tyr Trp Cys Asp Ala Arg Met Asp Lys Ile Glu Arg Ile Asp Leu Glu Thr Gly Glu Asn Arg Glu Val Val Leu Ser Ser Asn Asn Met Asp Met Phe Ser Val Ser Val Phe Glu Asp Phe Ile Tyr Trp Ser Asp Arg Thr His Ala Asn Gly Ser Ile Lys Arg Gly Cys Lys Asp Asn Ala Thr Asp Ser Val Pro Leu Arg Thr Gly Ile Gly Val Gln Leu Lys Asp Ile Lys Val Phe Asn Arg Asp Arg Gln Lys Gly Thr Asn Val Cys Ala Val Ala Asn Gly Gly Cys Gln Gln Leu Cys Leu Tyr Arg Gly Gly Gly Gln Arg Ala Cys Ala Cys Ala His Gly Met Leu Ala Glu Asp Gly Ala Ser Cys Arg Glu Tyr Ala Gly Tyr Leu Leu Tyr Ser Glu Arg Thr Ile Leu Lys Ser Ile His Leu Ser Asp Glu Arg Asn Leu Asn Ala Pro Val Gln Pro Phe Glu Asp Pro Glu His Met Lys Asn Val Ile Ala Leu Ala Phe Asp Tyr Arg Ala Gly Thr Ser Pro Gly Thr Pro Asn Arg Ile Phe Phe Ser Asp Ile His Phe Gly Asn Ile Gln Gln Ile Asn Asp Asp Gly Ser Gly Arg Thr Thr Ile Val Glu Asn Val Gly Ser Val Glu Gly -2280 Leu Ala Tyr His Arg Gly Trp Asp Thr Leu Tyr Trp Thr Ser Tyr Thr Thr Ser Thr Ile Thr Arg His Thr Val Asp Gln Thr Arg Pro Gly Ala Phe Glu Arg Glu Thr Val Ile Thr Met Ser Gly Asp Asp His Pro Arg Ala Phe Val Leu Asp Glu Cys Gln Asn Leu Met Phe Trp Thr Asn Trp

(Sh t 46 of 91)

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3	C)		2340		C	T1.			. 1 -		_		350		•
	2	355			Ser	2	2360				2	365			
	Leu 2370	Thr	Leu	Ile	Glu 2	Lys 2375	Asp	Ile	Arg		Pro 380	Asn	Gly	Leu	Ala
Ile 385	Asp	His	Arg		Glu 2390					Ser 2395	_		Thr		Asp 400
	Ile	Glu		Cys	Glu	Tyr		Gly			_	Tyr			Leu
Lvs	Ser	Glu			His		Phe			Ala		ጥህድ		2415 Glu	His
-		2	2420				2	425				2	2430		
	2	435			Trp	2	2440				2	445			
	Val 2450			Asp	Met 2		Leu		Arg		Asp 460	Ile	Pro	Gln	Gln
	Met	Gly	Ile		Ala	Val	Ala	Asn			Asn	Ser	Cys		
465	Dwa	Cuc	7.~~		2470	N cn	C1	C1		2475	N	T 0	Cuc		480
			2	2485	Asn			2	2490		_			2495	
		7	2500		Val		2	505				2	2510		
	2	2515			Cys	2	2520				2	2525			
2	2530			-		2535				2	2540				
Cys 545	Asp	Gly	Val		His 2550	Cys	Lys	Asp	_	Ser 2555	Asp	Glu	Lys		Ser 2560
	Cys	Asn			Arg	Cys	Lys	_		Phe		Gln	_		_
Gly	Arg		Val	Ser	Asn			Trp	Cys	Asn	Gly				Cys
Gly	-				Glu	Ile					Thr			Gly	Val
_			Arg	Cys	Arg			Ser	Cys				Ser	Ser	Arg
		Gln	Phe	Val	Asp		Glu	Asp	Ala			Glu	Met	Asn	Cys
625				:	2630				:	2635	_			7	2640
				2645				:	2650		-		_	2655	
			2660		Glu			2665		-	_	;	2670		
	_ ;	2675	_		Asn		2680	_	_	_	:	2685			
_	Pro 2690	_		Lys	Arg	Pro 2695		_	Pro		Asn 2700	Tyr	Phe	Ala	Cys
	Ser				Ile 2710		Met	Ser	_	Thr 2715	-	Asp	Lys		Asp 2720
		Glu			Glu		Glu			Cys		Lys		Cys 2735	Ser,
Glu	Ala			Glu		Gln			Arg		Ile			Gln	Trp
Leu		Asp	-	Ser	Asp			Gly	Asp	Gly			Glu		Ala
		Glu					Gly					Ser		Pro	Gly
Thr	His		Cys			Glu		Trp			Asp		Asp		Asp 2800
785 Cys		Asp		Ala	Asp		Ser		Thr	Ala		Cys	Lei	Tyr	Asn
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Ser Thr Cys Asp Asp Arg Glu Phe Met Cys Gln Asn Arg Leu Cys Ile Pro Lys His Phe Val Cys Asp His Asp Arg Asp Cys Ala Asp Gly Ser Asp Glu Ser Pro Glu Cys Glu Tyr Pro Thr Cys Gly Pro Asn Glu Phe Arg Cys Ala Asn Gly Arg Cys Leu Ser Ser Arg Gln Trp Glu Cys Asp Gly Glu Asn Asp Cys His Asp His Ser Asp Glu Ala Pro Lys Asn Pro His Cys Thr Ser Pro Glu His Lys Cys Asn Ala Ser Ser Gln Phe Leu .2910 Cys Ser Ser Gly Arg Cys Val Ala Glu Ala Leu Leu Cys Asn Gly Gln Asp Asp Cys Gly Asp Gly Ser Asp Glu Arg Gly Cys His Val Asn Glu Cys Leu Ser Arg Lys Leu Ser Gly Cys Ser Gln Asp Cys Glu Asp Leu Lys Ile Gly Phe Lys Cys Arg Cys Arg Pro Gly Phe Arg Leu Lys Asp Asp Gly Arg Thr Cys Ala Asp Leu Asp Glu Cys Ser Thr Thr Phe Pro Cys Ser Gln Leu Cys Ile Asn Thr His Gly Ser Tyr Lys Cys Leu Cys Val Glu Gly Tyr Ala Pro Arg Gly Gly Asp Pro His Ser Cys Lys Ala Val Thr Asp Glu Glu Pro Phe Leu Ile Phe Ala Asn Arg Tyr Tyr Leu Arg Lys Leu Asn Leu Asp Gly Ser Asn Tyr Thr Leu Leu Lys Gln Gly Leu Asn Asn Ala Val Ala Leu Ala Phe Asp Tyr Arg Glu Gln Met Ile Tyr Trp Thr Gly Val Thr Thr Gln Gly Ser Met Ile Arg Arg Met His Leu Asn Gly Ser Asn Val Gln Val Leu His Arg Thr Gly Leu Ser Asn Pro Asp Gly Leu Ala Val Asp Trp Val Gly Gly Asn Leu Tyr Trp Cys Asp Lys Gly Arg Asp Thr Ile Glu Val Ser Lys Leu Asn Gly Ala Tyr Arg Thr Val Leu Val Ser Ser Gly Leu Arg Glu Pro Arg Ala Leu Val Val Asp Val Gln Asn Gly Tyr Leu Tyr Trp Thr Asp Trp Gly Asp His Ser Leu Ile Gly Arg Ile Gly Met Asp Gly Ser Gly Arg Ser Ile Ile Val Asp Thr Lys Ile Thr Trp Pro Asn Gly Leu Thr Val Asp Tyr Val Thr Glu Arg Ile Tyr Trp Ala Asp Ala Arg Glu Asp Tyr Ile Glu Phe Ala Ser Leu Asp Gly Ser Asn Arg His Val Val Leu Ser Gln Asp Ile Pro His Ile Phe Ala Leu Thr Leu Phe Glu Asp Tyr Val Tyr Trp Thr 3245 . Asp Trp Glu Thr Lys Ser Ile Asn Arg Ala His Lys Thr Thr Gly Ala Asn Lys Thr Leu Leu Ile Ser Thr Leu His Arg Pro Met Asp Leu His Val Phe His Ala Leu Arg Gln Pro Asp Val Pro Asn His Pro Cys Lys

Val Asn Asn Gly Gly Cys Ser Asn Leu Cys Leu Leu Ser Pro Gly Gly . Gly His Lys Cys Ala Cys Pro Thr Asn Phe Tyr Leu Gly Gly Asp Gly Arg Thr Cys Val Ser Asn Cys Thr Ala Ser Gln Phe Val Cys Lys Asn Asp Lys Cys Ile Pro Phe Trp Trp Lys Cys Asp Thr Glu Asp Asp Cys Gly Asp His Ser Asp Glu Pro Pro Asp Cys Pro Glu Phe Lys Cys Arg Pro Gly Gln Phe Gln Cys Ser Thr Gly Ile Cys Thr Asn Pro Ala Phe Ile Cys Asp Gly Asp Asn Asp Cys Gln Asp Asn Ser Asp Glu Ala Asn Cys Asp Ile His Val Cys Leu Pro Ser Gln Phe Lys Cys Thr Asn Thr Asn Arg Cys Ile Pro Gly Ile Phe Arg Cys Asn Gly Gln Asp Asn Cys Gly Asp Glu Asp Glu Arg Asp Cys Pro Glu Val Thr Cys Ala Pro Asn Gln Phe Gln Cys Ser Ile Thr Lys Arg Cys Ile Pro Arg Val Trp Val Cys Asp Arg Asp Asn His Cys Val Asp Gly Ser Asp Glu Pro Ala Asn Cys Thr Gln Met Thr Cys Gly Val Asp Glu Phe Arg Cys Lys Asp Ser Gly Arg Cys Ile Pro Ala Arg Trp Lys Cys Asp Gly Glu Asp Asp Cys Gly Asp Gly Ser Asp Glu Pro Lys Glu Glu Cys Asp Glu Arg Thr Cys Glu Pro Tyr Gln Phe Arg Cys Lys Asn Asn Arg Cys Val Pro Gly Arg Trp Gln Cys Asp Tyr Asp Asn Asp Cys Gly Asp Asn Ser Asp Glu Glu Ser Cys Thr Pro Arg Pro Cys Ser Glu Ser Glu Phe Phe Cys Ala Asn Gly Arg Cys Ile Ala Gly Arg Trp Lys Cys Asp Gly Asp His Asp Cys Ala Asp Gly Ser Asp Glu Lys Asp Cys Thr Pro Arg Cys Asp Met Asp Gln Phe Gln Cys Lys Ser Gly His Cys Ile Pro Leu Arg Trp Pro Cys Asp Ala Asp Ala Asp Cys Met Asp Gly Ser Asp Glu Glu Ala Cys Gly Thr Gly Val Arg Thr Cys Pro Leu Asp Glu Phe Gln Cys Asn Asn Thr Leu Cys Lys Pro Leu Ala Trp Lys Cys Asp Gly Glu Asp Asp Cys Gly Asp Asn Ser Asp Glu Asn Pro Glu Glu Cys Ala Arg Phe Ile Cys Pro Pro Asn Arg Pro Phe Arg Cys Lys Asn Asp Arg Val Cys Leu Trp Ile Gly Arg Gln Cys Asp Gly Val Asp Asn Cys Gly Asp Gly Thr Asp Glu Glu Asp Cys Glu Pro Pro Thr Ala Gln Asn Pro His Cys Lys Asp Lys Lys Glu Phe Leu Cys Arg Asn Gln Arg Cys Leu Ser Ser Leu

(Sheet 49 of 91)

Arg Cys Asn Met Phe Asp Asp Cys Gly Asp Gly Ser Asp Glu Glu Asp Cys Ser Ile Asp Pro Lys Leu Thr Ser Cys Ala Thr Asn Ala Ser Met Cys Gly Asp Glu Ala Arg Cys Val Arg Thr Glu Lys Ala Ala Tyr Cys Ala Cys Arg Ser Gly Phe His Thr Val Pro Gly Gln Pro Gly Cys Gln Asp Ile Asn Glu Cys Leu Arg Phe Gly Thr Cys Ser Gln Leu Trp Asn Lys Pro Lys Gly Gly His Leu Cys Ser Cys Ala Arg Asn Phe Met Lys Thr His Asn Thr Cys Lys Ala Glu Gly Ser Glu Tyr Gln Val Leu Tyr Ile Ala Asp Asp Asn Glu Ile Arg Ser Leu Phe Pro Gly His Pro His Ser Ala Tyr Glu Gln Thr Phe Gln Gly Asp Glu Ser Val Arg Ile Asp Ala Met Asp Val His Val Lys Ala Gly Arg Val Tyr Trp Thr Asn Trp His Thr Gly Thr Ile Ser Tyr Arg Ser Leu Pro Pro Ala Ala Pro Pro Thr Thr Ser Asn Arg His Arg Arg Gln Ile Asp Arg Gly Val Thr His Leu Asn Ile Ser Gly Leu Lys Met Pro Arg Gly Ile Ala Ile Asp Trp Val Ala Gly Asn Val Tyr Trp Thr Asp Ser Gly Arg Asp Val Ile Glu Val Ala Gln Met Lys Gly Glu Asn Arg Lys Thr Leu Ile Ser Gly Met Ile Asp Glu Pro His Ala Ile Val Val Asp Pro Leu Arg Gly Thr Met . . Tyr Trp Ser Asp Trp Gly Asn His Pro Lys Ile Glu Thr Ala Ala Met Asp Gly Thr Leu Arg Glu Thr Leu Val Gln Asp Asn Ile Gln Trp Pro Thr Gly Leu Ala Val Asp Tyr His Asn Glu Arg Leu Tyr Trp Ala Asp Ala Lys Leu Ser Val Ile Gly Ser Ile Arg Leu Asn Gly Thr Asp Pro Ile Val Ala Ala Asp Ser Lys Arg Gly Leu Ser His Pro Phe Ser Ile Asp Val Phe Glu Asp Tyr Ile Tyr Gly Val Thr Tyr Ile Asn Asn Arg Val Phe Lys Ile His Lys Phe Gly His Ser Pro Leu Tyr Asn Leu Thr Gly Gly Leu Ser His Ala Ser Asp Val Val Leu Tyr His Gln His Lys Gln Pro Glu Val Thr Asn Pro Cys Asp Arg Lys Lys Cys Glu Trp Leu Cys Leu Leu Ser Pro Ser Gly Pro Val Cys Thr Cys Pro Asn Gly Lys Arg Leu Asp Asn Gly Thr Cys Val Pro Val Pro Ser Pro Thr Pro Pro Pro Asp Ala Pro Arg Pro Gly Thr Cys Thr Leu Gln Cys Phe Asn Gly Gly Ser Cys Phe Leu Asn Ala Arg Arg Gln Pro Lys Cys Arg Cys Gln Pro Arg Tyr Thr Gly Asp Lys Cys Glu Leu Asp Gln Cys Trp Glu Tyr

Cys His Asn Gly Gly Thr Cys Ala Ala Ser Pro Ser Gly Met Pro Thr Cys Arg Cys Pro Thr Gly Phe Thr Gly Pro Lys Cys Thr Ala Gln Val Cys Ala Gly Tyr Cys Ser Asn Asn Ser Thr Cys Thr Val Asn Gln Gly Asn Gln Pro Gln Cys Arg Cys Leu Pro Gly Phe Leu Gly Asp Arg Cys Gln Tyr Arg Gln Cys Ser Gly Phe Cys Glu Asn Phe Gly Thr Cys Gln Met Ala Ala Asp Gly Ser Arg Gln Cys Arg Cys Thr Val Tyr Phe Glu Gly Pro Arg Cys Glu Val Asn Lys Cys Ser Arg Cys Leu Gln Gly Ala Cys Val Val Asn Lys Gln Thr Gly Asp Val Thr Cys Asn Cys Thr Asp Gly Arg Val Ala Pro Ser Cys Leu Thr Cys Ile Asp His Cys Ser Asn Gly Gly Ser Cys Thr Met Asn Ser Lys Met Met Pro Glu Cys Gln Cys Pro Pro His Met Thr Gly Pro Arg Cys Gln Glu Gln Val Val Ser Gln Gln Gln Pro Gly His Met Ala Ser Ile Leu Ile Pro Leu Leu Leu Leu Leu Leu Leu Val Ala Gly Val Val Phe Trp Tyr Lys Arg Arg Val Arg Gly Ala Lys Gly Phe Gln His Gln Arg Met Thr Asn Gly Ala Met Asn Val Glu Ile Gly Asn Pro Thr Tyr Lys Met Tyr Glu Gly Gly Glu Pro Asp Asp Val Gly Gly Leu Leu Asp Ala Asp Phe Ala Leu Asp Pro Asp Lys Pro Thr Asn Phe Thr Asn Pro Val Tyr Ala Thr Leu Tyr Met Gly Gly His Gly Ser Arg His Ser Leu Ala Ser Thr Asp Glu Lys Arg Glu Leu Gly Arg Gly Pro Glu Asp Glu Ile Gly Asp Pro Leu Ala

GC	TAC	CAATC	C AI	CTGO	STCTC	стс	CAGC	TCC	TTCT	TTCI	GC A	AC F		GG F			5 5
L	AA ys 5	CTC Leu	CTT Leu	CAT His	CCA Pro	AGT Ser 10	CTG Leu	GTT Val	CTT Leu	CTC Leu	CTC Leu 15	TTG Leu	GTC Val	CTC Leu	CTG Leu	CCC Pro 20	103
A	CA hr	GAC Asp	GCC Ala	TCA Ser	GTC Val 25	TCT _. Ser	GGA Gly	AAA Lys	CCG Pro	CAG Gln 30	TAT Tyr	ATG Met	GTT Val	CTG Leu	GTC Val 35	CCC Pro	151
											GGC Gly						199
											TCC Ser						247
											GCG Ala						295
H											TCA Ser 95						343
											ACC Thr						391
											AGT Ser						439
											ACA Thr						487
											AAT Asn						535
V							••				ATC Ile 175						583
											TCT Ser						631
										Val	GTA Val						679
C	GGA	AGG	ACA	GAG	CAC	ССТ	TTC	ACC	GTG	GAG	GAA	ттт	GTT	СТТ	ccc	AAG	727

FIG. 13A

405

8449-134 (Sheet 52 of 91) Gly Arg Thr Glu His Pro Phe Thr Val Glu Glu Phe Val Leu Pro Lys 215 220 TTT GAA GTA CAA GTA ACA GTG CCA AAG ATA ATC ACC ATC TTG GAA GAA 775 Phe Glu Val Gln Val Thr Val Pro Lys Ile Ile Thr Ile Leu Glu Glu 235 240 GAG ATG AAT GTA TCA GTG TGT GGC CTA TAC ACA TAT GGG AAG CCT GTC Glu Met Asn Val Ser Val Cys Gly Leu Tyr Thr Tyr Gly Lys Pro Val 823 245 250 260 CCT GGA CAT GTG ACT GTG AGC ATT TGC AGA AAG TAT AGT GAC GCT TCC 871 Pro Gly His Val Thr Val Ser Ile Cys Arg Lys Tyr Ser Asp Ala Ser 265 GAC TGC CAC GGT GAA GAT TCA CAG GCT TTC TGT GAG AAA TTC AGT GGA 919 Asp Cys His Gly Glu Asp Ser Gln Ala Phe Cys Glu Lys Phe Ser Gly 280 285 290 CAG CTA AAC AGC CAT GGC TGC TTC TAT CAG CAA GTA AAA ACC AAG GTC 967 Gln Leu Asn Ser His Gly Cys Phe Tyr Gln Gln Val Lys Thr Lys Val 295 300 TTC CAG CTG AAG AGG AAG GAG TAT GAA ATG AAA CTT CAC ACT GAG GCC 1015 Phe Gln Leu Lys Arg Lys Glu Tyr Glu Met Lys Leu His Thr Glu Ala 310 315 CAG ATC CAA GAA GAA GGA ACA GTG GTG GAA TTG ACT GGA AGG CAG TCC 1063 Gln Ile Gln Glu Glu Gly Thr Val Val Glu Leu Thr Gly Arg Gln Ser 325 . 330 340 AGT GAA ATC ACA AGA ACC ATA ACC AAA CTC TCA TTT GTG AAA GTG GAC 1111 Ser Glu Ile Thr Arg Thr Ile Thr Lys Leu Ser Phe Val Lys Val Asp 345 350 TCA CAC TTT CGA CAG GGA ATT CCC TTC TTT GGG CAG GTG CGC CTA GTA 1159 Ser His Phe Arg Gln Gly Ile Pro Phe Phe Gly Gln Val Arg Leu Val 360 365 GAT GGG AAA GGC GTC CCT ATA CCA AAT AAA GTC ATA TTC ATC AGA GGA 1207 Asp Gly Lys Gly Val Pro Ile Pro Asn Lys Val Ile Phe Ile Arg Gly 375 AAT GAA GCA AAC TAT TAC TCC AAT GCT ACC ACG GAT GAG CAT GGC CTT 1255 Asn Glu Ala Asn Tyr Tyr Ser Asn Ala Thr Thr Asp Glu His Gly Leu 390 395

FIG. 13A

430

415

1303

1351

420

GTA CAG TTC TCT ATC AAC ACC ACC AAC GTT ATG GGT ACC TCT CTT ACT

Val Gln Phe Ser Ile Asn Thr Thr Asn Val Met Gly Thr Ser Leu Thr

GTT AGG GTC AAT TAC AAG GAT CGT AGT CCC TGT TAC GGC TAC CAG TGG

Val Arg Val Asn Tyr Lys Asp Arg Ser Pro Cys Tyr Gly Tyr Gln Trp

GTG TCA GAA GAA CAC GAA GAG GCA CAT CAC ACT GCT TAT CTT GTG TTC Val Ser Glu Glu His Glu Glu Ala His His Thr Ala Tyr Leu Val Phe

445

410

425

440

470

8449-134 (Sh t 53 of 91) TCC CCA AGC AAG AGC TTT GTC CAC CTT GAG CCC ATG TCT CAT GAA CTA Ser Pro Ser Lys Ser Phe Val His Leu Glu Pro Met Ser His Glu Leu 465 CCC TGT GGC CAT ACT CAG ACA GTC CAG GCA CAT TAT ATT CTG AAT GGA Pro Cys Gly His Thr Gln Thr Val Gln Ala His Tyr Ile Leu Asn Gly

GGC ACC CTG CTG GGG CTG AAG AAG CTC TCC TTT TAT TAT CTG ATA ATG
Gly Thr Leu Leu Gly Leu Lys Lys Leu Ser Phe Tyr Tyr Leu Ile Met
485
500

475

GCA AAG GGA GGC ATT GTC CGA ACT GGG ACT CAT GGA CTG CTT GTG AAG
Ala Lys Gly Gly Ile Val Arg Thr Gly Thr His Gly Leu Leu Val Lys
505
510
515

CAG GAA GAC ATG AAG GGC CAT TTT TCC ATC TCA ATC CCT GTG AAG TCA
Gln Glu Asp Met Lys Gly His Phe Ser Ile Ser Ile Pro Val Lys Ser
520 530

GAC ATT GCT CCT GTC GCT CGG TTG CTC ATC TAT GCT GTT TTA CCT ACC

Asp Ile Ala Pro Val Ala Arg Leu Leu Ile Tyr Ala Val Leu Pro Thr

535

540

545

GGG GAC GTG ATT GGG GAT TCT GCA AAA TAT GAT GTT GAA AAT TGT CTG
Gly Asp Val Ile Gly Asp Ser Ala Lys Tyr Asp Val Glu Asn Cys Leu
550
560

GCC AAC AAG GTG GAT TTG AGC TTC AGC CCA TCA CAA AGT CTC CCA GCC 1783
Ala Asn Lys Val Asp Leu Ser Phe Ser Pro Ser Gln Ser Leu Pro Ala
565 570 580

TCA CAC GCC CAC CTG CGA GTC ACA GCG GCT CCT CAG TCC GTC TGC GCC

Ser His Ala His Leu Arg Val Thr Ala Ala Pro Gln Ser Val Cys Ala

585

590

595

CTC CGT GCT GTG GAC CAA AGC GTG CTG CTC ATG AAG CCT GAT GCT GAG

Leu Arg Ala Val Asp Gln Ser Val Leu Leu Met Lys Pro Asp Ala Glu

600 605 610

CTC TCG GCG TCC TCG GTT TAC AAC CTG CTA CCA GAA AAG GAC CTC ACT 1927
Leu Ser Ala Ser Ser Val Tyr Asn Leu Leu Pro Glu Lys Asp Leu Thr
615 620 625

GGC TTC CCT GGG CCT TTG AAT GAC CAG GAC GAT GAA GAC TGC ATC AAT

Gly Phe Pro Gly Pro Leu Asn Asp Gln Asp Asp Glu Asp Cys Ile Asn

630

635

640

CGT CAT AAT GTC TAT ATT AAT GGA ATC ACA TAT ACT CCA GTA TCA AGT
Arg His Asn Val Tyr Ile Asn Gly Ile Thr Tyr Thr Pro Val Ser Ser
655 660

ACA AAT GAA AAG GAT ATG TAC AGC TTC CTA GAG GAC ATG GGC TTA AAG
Thr Asn Glu Lys Asp Met Tyr Ser Phe Leu Glu Asp Met Gly Leu Lys
665 670 675

GCA TTC ACC AAC TCA AAG ATT CGT AAA CCC AAA ATG TGT CCA CAG CTT 2119
Ala Phe Thr Asn Ser Lys Ile Arg Lys Pro Lys Met Cys Pro Gln Leu
680 685 690

FIG. 13A

84	449)-13	34	•									(5	h	t 54	of	91
CAA Gln	CAG Gln	TAT Tyr 695	GAA Glu	ATG Met	CAT His	GGA Gly	CCT Pro 700	GIU	GGT Gly	CTA	CGT Arg	GTA Val 705	Gly	TTT Phe	TAT Tyr	216	57
GAG Glu	TCA Ser 710	പാവ	GTA Val	ATG Met	GGA Gly	AGA Arg 715	GGC Gly	CAT His	GCA Ala	. CGC Arg	CTG Leu 720	Val	CAT His	GTT Val	GAA Glu	221	.5
GAG Glu 725	CCT Pro	CAC His	ACG Thr	GAG Glu	ACC Thr 730	vaı	CGA Arg	AAG Lys	TAC Tyr	TTC Phe 735	CCT Pro	GAG Glu	ACA Thr	TGG Trp	ATC Ile 740	226	3
пр	Asp	Leu	GTG Val	745	vai	Asn	Ser	Ala	Gly 750	Val	Ala	Glu	Val	Gly 755	Val	231	1
7112	V 61.	FIO	GAC Asp 760	1111	TIE	Inr	GIU	765	Lys	Ala	Gly	Ala	Phe 770	Cys	Leu	235	9 .
TCT Ser	GAA Glu	GAT Asp 775	GCT Ala	GGA Gly	CTT Leu	GGT Gly	ATC Ile 780	TCT Ser	TCC Ser	ACT Thr	GCC Ala	TCT Ser 785	CTC Leu	CGA Arg	GCC Ala	240	7
rne	790	PIO	TTC Phe	Pne	vaı	795	Leu	Thr	Met	Pro	Tyr 800	Ser	Val	Ile	Arg	245	5
805	Ģīu	NIG	TTC Phe	mr	810	ьys	Ala	Thr	Val	Leu 815	Asn	Tyr	Leu	Pro	Lys 820	250	3
TGC Cys	ATC Ile	CGG Arg	GTC Val	AGT Ser 825	GTG Val	CAG Gln	CTG Leu	GAA Glu	GCC Ala 830	TCT Ser	CCC Pro	GCC Ala	TTC Phe	CTT Leu 835	GCT Ala	255	1
GTC Val	CCA Pro	GTG Val	GAG Glu 840	AAG Lys	GAA Glu	CAA Gln	GCG Ala	CCT Pro 845	CAC His	TGC Cys	ATC Ile	TGT Cys	GCA Ala 850	AAC Asn	GGG Gly	259	9.
CGG Arg	CAA Gln	ACT Thr 855	GTG Val	TCC Ser	TGG Trp	Ala	GTA Val 860	ACC Thr	CCA Pro	AAG Lys	TCA Ser	TTA Leu 865	GGA Gly	TAA naA	GTG Val	264	7
AAT Asn	TȚC Phe 870	ACT Thr	GTG Val	AGC Ser	GCA Ala	GAG Glu 875	GCA Ala	CTA Leu	GAG Glu	TCT Ser	CAA Gln 880	GAG Glu	CTG Leu	TGT ·	GGG Gly	269	5
ACT Thr 885	GAG Glu	GTG Val	CCT Pro	ser	GTT Val 890	CCT Pro	GAA Glu	CAC His	GGA Gly	AGG Arg 895	AAA Lys	GAC Asp	ACA Thr	GTC Val	ATC Ile 900	2743	3
AAG Lys	CCT Pro	CTG Leu	TTG Leu	GTT Val 905	GAA Glu	CCT Pro	GAA Glu	GGA Gly	CTA Leu 910	GAG Glu	AAG Lys	GAA Glu	ACA Thr	ACA Thr 915	TTC Phe	2791	l
AAC Asn	TCC Ser	CTA Leu	CTT Leu 920	TGT Cys	CCA Pro	TCA Ser	Gly	GGT Gly 925	GAG Glu	GTT Val	TCT Ser	GAA Glu	GAA Glu 930	TTA Leu	TCC Ser	2839	9

FIG. 13A

8449-134 (She t 55 of 91) CTG AAA CTG CCA CCA AAT GTG GTA GAA GAA TCT GCC CGA GCT TCT GTC 2887 Leu Lys Leu Pro Pro Asn Val Val Glu Glu Ser Ala Arg Ala Ser Val 935 940 TCA GTT TTG GGA GAC ATA TTA GGC TCT GCC ATG CAA AAC ACA CAA AAT 2935 Ser Val Leu Gly Asp Ile Leu Gly Ser Ala Met Gln Asn Thr Gln Asn 950 955 CTT CTC CAG ATG CCC TAT GGC TGT GGA GAG CAG AAT ATG GTC CTC TTT 2983 Leu Leu Gln Met Pro Tyr Gly Cys Gly Glu Gln Asn Met Val Leu Phe 965 975 980 GCT CCT AAC ATC TAT GTA CTG GAT TAT CTA AAT GAA ACA CAG CAG CTT 3031 Ala Pro Asn Ile Tyr Val Leu Asp Tyr Leu Asn Glu Thr Gln Gln Leu 985 990 ACT CCA GAG GTC AAG TCC AAG GCC ATT GGC TAT CTC AAC ACT GGT TAC 3079 Thr Pro Glu Val Lys Ser Lys Ala Ile Gly Tyr Leu Asn Thr Gly Tyr 1000 1005 CAG AGA CAG TTG AAC TAC AAA CAC TAT GAT GGC TCC TAC AGC ACC TTT 3127 Gln Arg Gln Leu Asn Tyr Lys His Tyr Asp Gly Ser Tyr Ser Thr Phe 1015 1020 GGG GAG CGA TAT GGC AGG AAC CAG GGC AAC ACC TGG CTC ACA GCC TTT 3175 Gly Glu Arg Tyr Gly Arg Asn Gln Gly Asn Thr Trp Leu Thr Ala Phe 1030 1035 1040 GTT CTG AAG ACT TTT GCC CAA GCT CGA GCC TAC ATC TTC ATC GAT GAA 3223 Val Leu Lys Thr Phe Ala Gln Ala Arg Ala Tyr Ile Phe Ile Asp Glu 1050 1055 GCA CAC ATT ACC CAA GCC CTC ATA TGG CTC TCC CAG AGG CAG AAG GAC 3271 Ala His Ile Thr Gln Ala Leu Ile Trp Leu Ser Gln Arg Gln Lys Asp 1065 1070 1075 AAT GGC TGT TTC AGG AGC TCT GGG TCA CTG CTC AAC AAT GCC ATA AAG 3319 Asn Gly Cys Phe Arg Ser Ser Gly Ser Leu Leu Asn Asn Ala Ile Lys 1080 1085 GGA GGA GTA GAA GTG ACC CTC TCC GCC TAT ATC ACC ATC GCC 3367 Gly Gly Val Glu Asp Glu Val Thr Leu Ser Ala Tyr Ile Thr Ile Ala 1095 1100 1105 CTT CTG GAG ATT CCT CTC ACA GTC ACT CAC CCT GTT GTC CGC AAT GCC 3415 Leu Leu Glu Ile Pro Leu Thr Val Thr His Pro Val Val Arg Asn Ala 1110 1115 CTG TTT TGC CTG GAG TCA GCC TGG AAG ACA GCA CAA GAA GGG GAC CAT 3463 Leu Phe Cys Leu Glu Ser Ala Trp Lys Thr Ala Gln Glu Gly Asp His 1125 1130 1135 1140

FIG. 13A

1150

3511

3559

1155

1170

GGC AGC CAT GTA TAT ACC AAA GCA CTG CTG GCC TAT GCT TTT GCC CTG

Gly Ser His Val Tyr Thr Lys Ala Leu Leu Ala Tyr Ala Phe Ala Leu

GCA GGT AAC CAG GAC AAG AGG AAG GAA GTA CTC AAG TCA CTT AAT GAG

Ala Gly Asn Gln Asp Lys Arg Lys Glu Val Leu Lys Ser Leu Asn Glu

1165

1145

1160

(She t 56 of 91)

	GCT Ala 1					Asn					Glu					3607
Pro	AAG Lys 190				Gly					Pro						3655
	GTG Val			Thr					Leu					Ala		3703
	GCC Ala		Thr					Thr					Ile			3751
	ATC Ile	Thr					Ala					Ser				3799
	ACA Thr					His					Tyr					3847
Phe	ACC Thr 1270	Arg			Lys					Thr						3895
	TTT Phe			Lys					Asn					Leu		3943
	CAG Gln		Ser					Pro					Met			3991
	GGA Gly	Glu					Leu					Lys				4039
	CCA Pro					Phe					Gly					4087
Pro	CAA Gln 1350				Glu					Thr						4135
	AGT Ser			Tyr					Ser					Ala	_	4183
	GAT Asp		Lys					Phe		Pro					Val	4231
		Leu		Arg			His		Ser			Glu		Ser	AGC Ser	4279

(Sheet 57 of 91)

AAC Asn	His	GTC Val 1415	TTG Leu	ATT Ile	TAC Tyr	Leu	GAT Asp 1420	AAG Lys	GTG Val	TCA Ser	Asn	CAG Gln 1425	ACA Thr	CTG Leu	AGC Ser	4327
Leu	TTC Phe 430	TTC Phe	ACG Thr	GTT Val	CTG Leu	CAA Gln 1435	Asp	GTC Val	CCA Pro	Val	AGA Arg 1440	GAT Aşp	CTC Leu	AAA Lys	CCA Pro	4375
				Val	TAT Tyr 1450				Glu					Ala		4423
GCT Ala	GAG Glu	TAC Tyr	Asn	GCT Ala 1465	CCT Pro	TGC Cys	AGC Ser	Lys	GAT Asp 1470	CTT Leu	GGA Gly	AAT Asn	GCT Ala	TGAA 1	AGACCA	4474
					ro di ra di							CACAC	GAA (GACAC	CGTGTT	4534 4577

Ser Val Ser Gly Lys Pro Gln Tyr Met Val Leu Val Pro Ser Leu Leu His Thr Glu Thr Thr Glu Lys Gly Cys Val Leu Leu Ser Tyr Leu Asn Glu Thr Val Thr Val Ser Ala Ser Leu Glu Ser Val Arg Gly Asn Arg Ser Leu Phe Thr Asp Leu Glu Ala Glu Asn Asp Val Leu His Cys Val 55 Ala Phe Ala Val Pro Lys Ser Ser Ser Asn Glu Glu Val Met Phe Leu Thr Val Gln Val Lys Gly Pro Thr Gln Glu Phe Lys Lys Arg Thr Thr Val Met Val Lys Asn Glu Asp Ser Leu Val Phe Val Gln Thr Asp Lys 105 Ser Ile Tyr Lys Pro Gly Gln Thr Val Lys Phe Arg Val Val Ser Met 120 Asp Glu Asn Phe His Pro Leu Asn Glu Leu Ile Pro Leu Val Tyr Ile 135 Gln Asp Pro Lys Gly Asn Arg Ile Ala Gln Trp Gln Ser Phe Gln Leu 155 150 Glu Gly Gly Leu Lys Gln Phe Ser Phe Pro Leu Ser Ser Glu Pro Phe 170 Gln Gly Ser Tyr Lys Val Val Val Gln Lys Lys Ser Gly Gly Arg Thr 185 Glu His Pro Phe Thr Val Glu Glu Phe Val Leu Pro Lys Phe Glu Val 200 Gln Val Thr Val Pro Lys Ile Ile Thr Ile Leu Glu Glu Met Asn 220 Val Ser Val Cys Gly Leu Tyr Thr Tyr Gly Lys Pro Val Pro Gly His 235 230 Val Thr Val Ser Ile Cys Arg Lys Tyr Ser Asp Ala Ser Asp Cys His 245 Gly Glu Asp Ser Gln Ala Phe Cys Glu Lys Phe Ser Gly Gln Leu Asn 265 260 Ser His Gly Cys Phe Tyr Gln Gln Val Lys Thr Lys Val Phe Gln Leu 280 Lys Arg Lys Glu Tyr Glu Met Lys Leu His Thr Glu Ala Gln Ile Gln 295 Glu Glu Gly Thr Val Val Glu Leu Thr Gly Arg Gln Ser Ser Glu Ile 315 310 Thr Arg Thr Ile Thr Lys Leu Ser Phe Val Lys Val Asp Ser His Phe 330 Arg Gln Gly Ile Pro Phe Phe Gly Gln Val Arg Leu Val Asp Gly Lys 345 340 Gly Val Pro Ile Pro Asn Lys Val Ile Phe Ile Arg Gly Asn Glu Ala 360 Asn Tyr Tyr Ser Asn Ala Thr Thr Asp Glu His Gly Leu Val Gln Phe 380 375 Ser Ile Asn Thr Thr Asn Val Met Gly Thr Ser Leu Thr Val Arg Val 395 390 Asn Tyr Lys Asp Arg Ser Pro Cys Tyr Gly Tyr Gln Trp Val Ser Glu .410 Glu His Glu Glu Ala His His Thr Ala Tyr Leu Val Phe Ser Pro Ser 425 Lys Ser Phe Val His Leu Glu Pro Met Ser His Glu Leu Pro Cys Gly 440 His Thr Gln Thr Val Gln Ala His Tyr Ile Leu Asn Gly Gly Thr Leu 460 455 Leu Gly Leu Lys Lys Leu Ser Phe Tyr Tyr Leu Ile Met Ala Lys Gly

	•												•		
465					470					475					480
Gly				485	Gly				490	Leu			Gln	Glu 495	Asp
Met	Lys	Gly	His 500	Phe	Ser	Ile	Ser	11e 505	Pro	Val	Lys	Ser	Asp 510	Ile	Ala
Pro	Val	Ala 515	Arg	Leu	Leu	Ile	Tyr 520	Ala	Val	Leu	Pro	Thr 525	Gly	Asp	Val
Ile	Gly 530	Asp	Ser	Ala	Lys	Tyr 535	Asp	Val.	Glu	Asn _.	Cys 540	Leu	Ala	Asn	Lys
Val 545	Asp	Leu	Ser	Phe	Ser 550	Pro	Ser	Gln	Ser	Leu 555	Pro	Ala	Ser	His	Ala 560
				565					570		_		Leu	575	
	_		580					585		_			Leu 590		
		595	,				600			_		605	Gly ·		
_	610			_		615	_		_	_	620		Arg		
625	-			_	630					635			Thr		640
-	_	•	_	645				_	650	_			Ala	655	
		_	660					665					Gln 670		
		67 5	-				680	_		_		685	Glu		
	690					695					700		Glu Trp		
705	•			_	710					715	-		Thr		720
				725					730		-		Ser	735	
_			740		W			745					750 Phe		
	_	755	-				760					765	Gly		
	770					775					780		Cys		
785			_		790					795			Val		800
				805					810				Arg	815	
	_		820					825					830 Asn		
•	•	835					840					845			
	850					855					860		Lys		Leu
865					870					87 5			Asn		880
				885	• /				890	•			Leu	895	
	_		900	}	_			905	•			·	910 Ser		
		915					920)				925)		Gln
Gly	930			. 023	-	935					940				
		-			!		F	IG	•	13B	3		,		

Met Pro Tyr Gly Cys Gly Glu Gln Asn Met Val Leu Phe Ala Pro Asn Ile Tyr Val Leu Asp Tyr Leu Asn Glu Thr Gln Gln Leu Thr Pro Glu Val Lys Ser Lys Ala Ile Gly Tyr Leu Asn Thr Gly Tyr Gln Arg Gln Leu Asn Tyr Lys His Tyr Asp Gly Ser Tyr Ser Thr Phe Gly Glu Arg 1000 1005 Tyr Gly Arg Asn Gln Gly Asn Thr Trp Leu Thr Ala Phe Val Leu Lys Thr Phe Ala Gln Ala Arg Ala Tyr Ile Phe Ile Asp Glu Ala His Ile Thr Gln Ala Leu Ile Trp Leu Ser Gln Arg Gln Lys Asp Asn Gly Cys Phe Arg Ser Ser Gly Ser Leu Leu Asn Asn Ala Ile Lys Gly Gly Val Glu Asp Glu Val Thr Leu Ser Ala Tyr Ile Thr Ile Ala Leu Leu Glu Ile Pro Leu Thr Val Thr His Pro Val Val Arg Asn Ala Leu Phe Cys Leu Glu Ser Ala Trp Lys Thr Ala Gln Glu Gly Asp His Gly Ser His Val Tyr Thr Lys Ala Leu Leu Ala Tyr Ala Phe Ala Leu Ala Gly Asn Gln Asp Lys Arg Lys Glu Val Leu Lys Ser Leu Asn Glu Glu Ala Val Lys Lys Asp Asn Ser Val His Trp Glu Arg Pro Gln Lys Pro Lys Ala Pro Val Gly His Phe Tyr Glu Pro Gln Ala Pro Ser Ala Glu Val Glu Met Thr Ser Tyr Val Leu Leu Ala Tyr Leu Thr Ala Gln Pro Ala Pro Thr Ser Glu Asp Leu Thr Ser Ala Thr Asn Ile Val Lys Trp Ile Thr Lys Gln Gln Asn Ala Gln Gly Gly Phe Ser Ser Thr Gln Asp Thr Val Val Ala Leu His Ala Leu Ser Lys Tyr Gly Ala Ala Thr Phe Thr Arg Thr Gly Lys Ala Ala Gln Val Thr Ile Gln Ser Ser Gly Thr Phe Ser Ser Lys Phe Gln Val Asp Asn Asn Asn Arg Leu Leu Gln Gln Val Ser Leu Pro Glu Leu Pro Gly Glu Tyr Ser Met Lys Val Thr Gly Glu Gly Cys Val Tyr Leu Gln Thr Ser Leu Lys Tyr Asn Ile Leu Pro Glu Lys Glu Glu Phe Pro Phe Ala Leu Gly Val Gln Thr Leu Pro Gln Thr Cys Asp Glu Pro Lys Ala His Thr Sen Phe Gln Tle Ser Leu Ser Wal Ser Tyr Thr Gly Ser Arg Ser Ala Ser Asn Met Ala Ile Val Asp Val Lys Met Val Ser Gly Phe Ile Pro Leu Lys Pro Thr Glu Arg Ser Asn His Val Ser Arg Thr Glu Val Ser Ser Asn His Val Ile Tyr Leu Asp Lys Val Ser Asn Gln Thr Leu Ser Leu Phe Phe Thr Val Leu Gln Asp Val Pro Val Arg Asp Leu Lys Pro Ala Ile Val 1410 1420
Lys Val Tyr Asp Tyr Tyr Glu Thr Asp Glu Phe Ala Ile Ala Glu Tyr 425 1430 1435 1440
Asn Ala Pro Cys Ser Lys Asp Leu Gly Asn Ala 120

(She t 62 of 91)

CAGCGGTGCG AGCTCCAGGC CCATGCACTG AGGAGGCGGA AACAAGGGGA GCCCCAGAG CTCCATCAAG CCCCCTCCAA AGGCTCCCCT ACCCGGTCCA CGCCCCCCAC CCCCCCTCCC CGCCTCCTCC CAATTGTGCA TTTTTGCAGC CGGAGGCGGC TCCGAGATGG GGCTGTGAGC TTCGCCCGGG GAGGGGGAAA GAGCAGCGAG GAGTGAAGCG GGGGGGTGGG GTGAAGGGTT TGGATTTCGG GGCAGGGGC GCACCCCCGT CAGCAGGCCC TCCCCAAGGG GCTCGGAACT CTACCTCTTC ACCCACGCCC CTGGTGCGCT TTGCCGAAGG AAAGAATAAG AACAGAGAAG GAGGAGGGG AAAGGAGGAA AAGGGGGACC CCCCAACTGG GGGGGTGAA GGAGAGAAGT AGCAGGACCA GAGGGAAGG GGCTGCTGCT TGCATCAGCC CACACC ATG CTG ACC Met Leu Thr 1	60 120 180 240 300 360 420 475
CCG CCG TTG CTC CTG CTG CCC CTG CTC TCA GCT CTG GTC GCG GCG Pro Pro Leu Leu Leu Leu Pro Leu Leu Ser Ala Leu Val Ala Ala 5 10 15	523
GCT ATC GAC GCC CCT AAG ACT TGC AGC CCC AAG CAG TTT GCC TGC AGA Ala Ile Asp Ala Pro Lys Thr Cys Ser Pro Lys Gln Phe Ala Cys Arg 20 25 30 35	571
GAT CAA ATA ACC TGT ATC TCA AAG GGC TGG CGG TGC GAC GGT GAG AGG Asp Gln Ile Thr Cys Ile Ser Lys Gly Trp Arg Cys Asp Gly Glu Arg 40 45 50	619
GAC TGC CCA GAC GGA TCT GAC GAG GCC CCT GAG ATT TGT CCA CAG AGT Asp Cys Pro Asp Gly Ser Asp Glu Ala Pro Glu Ile Cys Pro Gln Ser 55 60 65	667
AAG GCC CAG CGA TGC CAG CCA AAC GAG CAT AAC TGC CTG GGT ACT GAG Lys Ala Gln Arg Cys Gln Pro Asn Glu His Asn Cys Leu Gly Thr Glu 70 75 80	715
CTG TGT GTT CCC ATG TCC CGC CTC TGC AAT GGG GTC CAG GAC TGC ATG Leu Cys Val Pro Met Ser Arg Leu Cys Asn Gly Val Gln Asp Cys Met 85 90 95	763
GAC GGC TCA GAT GAG GGG CCC CAC TGC CGA GAG CTC CAA GGC AAC TGC Asp Gly Ser Asp Glu Gly Pro His Cys Arg Glu Leu Gln Gly Asn Cys 100 115	811 .
TCT CGC CTG GGC TGC CAG CAC CAT TGT GTC CCC ACA CTC GAT GGG CCC Ser Arg Leu Gly Cys Gln His His Cys Val Pro Thr Leu Asp Gly Pro 120 125 130	859 '
ACC TGC TAC TGC AAC AGC AGC TTT CAG CTT CAG GCA GAT GGC AAG ACC Thr Cys Tyr Cys Asn Ser Ser Phe Gln Leu Gln Ala Asp Gly Lys Thr 135 140 145	907
TGC AAA GAT TTT GAT GAG TGC TCA GTG TAC GGC ACC TGC AGC CAG CTA Cys Lys Asp Phe Asp Glu Cys Ser Val Tyr Gly Thr Cys Ser Gln Leu 150 160	955
TGC ACC AAC ACA GAC GGC TCC TTC ATA TGT GGC TGT GTT GAA GGA TAC Cys Thr Asn Thr Asp Gly Ser Phe Ile Cys Gly Cys Val Glu Gly Tyr 165	1003
CTC CTG CAG CCG GAT AAC CGC TCC TGC AAG GCC AAG AAC GAG CCA GTA Leu Leu Gln Pro Asp Asn Arg Ser Cys Lys Ala Lys Asn Glu Pro Val 180 185 190 195	1051

FIG. 14A

(Sheet 63 of 91)

GAC Asp	CGG Arg	CCC Pro	CCT Pro	GTG Val 200	CTG Leu	TTG Leu	ATA Ile	GCC Ala	AAC Asn 205	TCC Ser	CAG Gln	AAC Asn	ATC Ile	TTG Leu 210	GCC Ala	1099
ACG Thr	TAC Tyr	CTG Leu	AGT Ser 215	GGG Gly	GCC Ala	CAG Gln	GTG Val	TCT Ser 220	ACC Thr	ATC Ile	ACA Thr	CCT Pro	ACG Thr 225	AGC Ser	ACG Thr	1147
CGG Arg	CAG Gln	ACC Thr 230	ACA Thr	GCC Ala	ATG Met	GAC Asp	TTC Phe 235	AGC Ser	TAT Tyr	GCC Ala	AAC Asn	GAG Glu 240	ACC Thr	GTA Val	TGC Cys	1195
TGG Trp	GTG Val 245	CAT His	GTT Val	GGG Gly	GAC Asp	AGT Ser 250	GCT Ala	GCT Ala	CAG Gln	ACG Thr	CAG Gln 255	CTC Leu	AAG Lys	TGT Cys	GCC Ala	1243
CGC Arg 260	ATG Met	CCT Pro	GGC Gly	CTA Leu	AAG Lys 265	GGC Gly	TTC Phe	GTG Val	GAT Asp	GAG Glu 270	CAC His	ACC Thr	ATC Ile	AAC Asn	ATC Ile 275	1291
												GAC Asp				1339
												ATC Ile				1387
												CTG Leu 320				1435
												AAG Lys				1483
									Glu			GAC Asp				1531
												TTT Phe			Gly ·	1579
												GCA Ala		Ala		1627
												GGC Gly 400				1675
												CTG Leu				1723
	Asn										Ala	AAT Asn			CAG Gln 435	1771

FIG. 14A

(Sh et 64 of 91)

												ACC Thr				1819
												ATC Ile				1867
AGG Arg	CGT Arg	CAG Gln 470	CCC Pro	CGA Arg	GTĞ Val	AGG Arg	AGC Ser 475	CÀT His	GCC Ala	TGT Cys	GAA Glu	AAC Asn 480	GAC Asp	CAG Gln	TAT Tyr	1915
												GCC Ala				1963
												CTG Leu				2011
GGG Gly	AAG Lys	TCA Ser	TGC Cys	AAG Lys 520	AAG Lys	CCG Pro	GAG Glu	CAT His	GAG Glu 525	CTG Leu	TTC Phe	CTC Leu	GTG Val	TAT Tyr 530	GGC Gly	2059
									Met			GGG Gly				2107
CCG Pro	GAT Asp	GAG Glu 550	His	ATG Met	ATC Ile	CCC Pro	ATT Ile 555	GAA Glu	AAC Asn	CTC Leu	ATG Met	AAC Asn 560	CCC Pro	CGA Arg	GCC Ala	2155
CTG Leu	GAC Asp 565	Phe	CAC His	GCT Ala	GAG Glu	ACC Thr 570	GC	TTC Phe	ATC Ile	TAC Tyr	TTT Phe 575	GCC Ala	GAC Asp	ACC Thr	ACC Thr	2203
AGC Ser 580	Tyr	CTC Leu	ATT Ile	GGC	CGC Arg 585	Gln	AAG Lys	ATT	GAT Asp	GGC Gly 590	ACT Thr	GAG Glu	CGG Arg	GAG Glu	ACC Thr 595	2251
Ile	Leu	Lys	Asp	Gly 600	Ile	His	Asn	Val	. Glu 605	Gly	Val	GCC Ala	Val	Asp 610	Trp	2299
Met	Gly	Asp	Asn 615	Leu	Tyr	Trp	Thr	Asp 620	Asp	Gly	Pro	AAA Lys	Lys 625	Thr	Ile	2347
AGC Ser	GTC Val	GCC Ala 630	Arg	CTG Leu	GAG Glu	AAA Lys	GCT Ala 635	Ala	r CAG a Glr	ACC Thr	CGC Arg	Lys 640	Thr	· TTA	ATC Ile	2395
Glt	Gly 645	y Lys	s Met	Thr	. His	650	Arç	y Ala	a Il€	e Val	. Val 655	L Asp) Pro	Leu	AAT Asn	2443
GG(G1 ₂ 660	y Tr	G ATO	TAC	TGC Trg	ACA Thi	c Asp	TG(G GA	G GAO u Glu	G GAC L Asp 670	o Pro	C AAC b Lys	GAC S Asp	C AG1	CGG Arg 675	2491

FIG. 14A

(Sheet 65 of 91)

						GCG Ala							Arg		_	2539
						GTG Val										2587
ATC Ile	CCG Pro	GCT Ala 710	GGG Gly	CGC Arg	CTC Leu	TAC Tyr	TGG Trp 715	GTG Val	GAT Asp	GCC Ala	TTC Phe	TAC Tyr 720	GAC Asp	CGC Arg	ATC Ile	2635
						GGC Gly 730										2683
						TTT Phe										2731
						AGT Ser										2779
GTA Val	GGA Gly	GGC Gly	GCA Ala 775	CCC Pro	CCC Pro	ACT Thr	GTG Val	ACC Thr 780	CTT Leu	CTG Leu	CGC Arg	AGT Ser	GAG Glu 785	CGG Arg	CCC Pro	2827
CCC Pro	ATC Ile	TTT Phe 790	GAG Glu	ATC Ile	CGA Arg	ATG Met	TAT Tyr 795	GAT Asp	GCC Ala	CAG Gln	CAG Gln	CAG Gln 800	CAA Gln	GTT Val	GCC	2875
ACC Thr	AAC Asn 805	AAA Lys	TGC Cys	CGG Arg	GTG Val	AAC Asn 810	AAT Asn	GGC Gly	GGC	TGC Cy s	AGC Ser 815	AGC Ser	CTG Leu	TGC Cys	TTG Leu	2923
GCC Ala 820	Thr	CCT Pro	GGG Gly	AGC Ser	CGC Arg 825	CAG Gln	TGC Cys	GCC Ala	TGT Cys	GCT Ala 830	GAG Glu	GAC Asp	CAG Gln	GTG Val	TTG Leu 835	2971 .
GAC Asp	GCA Ala	GAC Asp	GGC Gly	GTC Val 840	Thr	TGC Cys	TTG Leu	GCG Ala	AAC Asn 845	Pro	TCC Ser	TAC Tyr	GTG Val	CCT Pro 850	CCA Pro	3019
CCC Pro	CAG Gln	TGC Cys	Gln 855	Pro	GGC Gly	GAG Glu	TTT Phe	GCC Ala 860	Cys	GCC	AAC Asn	AGC Ser	CGC Arg 865	Cys	: ATC	3067
CAG Gln	GAG Glu	CGC Arg 870	Trp	AAG Lys	TGT Cys	GAC Asp	GGA Gly 875	/ Asp	AAC Asn	GAT Asp	TGC Cys	CTG Lev 880	. Asp	AAC Asr	AGT Ser	3115
GAT Asp	GAG Glu 885	ı Ala	CCF Pro	A GCC	CTC Leu	TGC Cys 890	His	CAC Glr	G CAC	ACC Thr	TGC Cys 895	Pro	TCG Ser	GAC Asp	C CGA	3163
TT(Phe 90(E Lys	TGC Cys	C GAG	AA E ISA L	AAC ASI 909	Arg	TG(Cy:	C ATO	C CCC	AAC Asi 910	n Arg	TG(CTO Lev	TGC Cy:	GAC S Asp 915	3211

FIG. 14A

(Sheet 66 of 91) 8449-134

GGG Gly	GAC Asp	AAT Asn	GAC Asp	TGT Cys 920	GGG Gly	AAC Asn	AGT Ser	GAA Glu	GAT Asp 925	GAG Glu	TCC Ser	AAT Asn	GCC Ala	ACT Thr 930	TGT Cys	3259
TCA Ser	GCC Ala	CGC Arg	ACC Thr 935	TGC Cys	CCC Pro	CCC Pro	AAC Asn	CAG Gln 940	TTC Phe	TCC Ser	TGT Cys	GCC Ala	AGT Ser 945	GGC	CGC Arg	3307
TGC Cys	ATC Ile	CCC Pro 950	ATC Ile	TCC Ser	TGG Trp	ACG Thr	TGT Cys 955	GAT Asp	CTG Leu	GAT Asp	GAC Asp	GAC Asp 960	TGT Cys	GGG	GAC Asp	3355
CGC Arg	TCT Ser 965	GAT Asp	GAG Glu	TCT Ser	GCT Ala	TCG Ser 970	TGT Cys	GCC Ala	TAT Tyr	CCC Pro	ACC Thr 975	TGC Cys	TTC Phe	CCC Pro	CTG Leu	3403
ACT Thr 980	CAG Gln	TTT Phe	ACC Thr	TGC Cys	AAC Asn 985	AAT Asn	GGC Gly	AGA Arg	TGT Cys	ATC Ile 990	AAC Asn	ATC Ile	AAC Asn	TGG Trp	AGA Arg 995	3451
TGC Cys	GAC Asp	AAT Asn	Asp	TAA neA 000	GAC Asp	TGT Cys	GGG Gly	Asp	AAC Asn .005	AGT Ser	GAC Asp	GAA Glu	Ala	GGC Gly 1010	TGC Cys	3499
AGC Ser	CAC His	Ser	TGT Cys 1015	TCT Ser	AGC Ser	ACC Thr	Gln	TTC Phe 020	AAG Lys	TGC Cys	AAC Asn	Ser	GGG Gly LO25	CGT Arg	TGC Cys	3547
ATC Ile	Pro	GAG Glu 1030	CAC His	TGG Trp	ACC Thr	Cys	GAT Asp 1035	GGG Gly	GAC Asp	AAT Asn	Asp	TGC Cys L040	GGA Gly	GAC Asp	TAC Tyr	3595
Ser	GAT Asp .045	GAG Glu	ACA Thr	CAC His	GCC Ala	AAC Asn 1050	TGC Cys	ACC Thr	AAC Asn	Gln	GCC Ala 1055	ACG Thr	AGG Arg	CCC Pro	CCT Pro	3643
				Thr	GAT Asp 1065				Cys					Leu		3691
			Arg		CGC Arg			Gly					Met			3739
		Glu			TGT Cys		Gly					Cys				3787
	Lys				AAG Lys	Asp					Ile					3835
Val					AAT Asn					Asn						3883
				Ala	TGC Cys 1145				Ser		Pro			Asn		3931
			•• .				FI	G.	14	A						

(Sheet 67 of 91)

									GAC Asp	
	_	1160			1165		•	1170		

TGT GGC GAC GGC TCA GAT GAG GGC GAG CTC TGC GAC CAG TGC TCT CTG

Cys Gly Asp Gly Ser Asp Glu Gly Glu Leu Cys Asp Gln Cys Ser Leu

1175

1180

1185

AAT AAC GGT GGC TGC AGC CAC AAC TGC TCA GTG GCA CCT GGC GAA GGC

Asn Asn Gly Gly Cys Ser His Asn Cys Ser Val Ala Pro Gly Glu Gly

1190 1195 1200

ATT GTG TGT TCC TGC CCT CTG GGC ATG GAG CTG GGG CCC GAC AAC CAC

11e Val Cys Ser Cys Pro Leu Gly Met Glu Leu Gly Pro Asp Asn His

1205

1210

1215

ACC TGC CAG ATC CAG AGC TAC TGT GCC AAG CAT CTC AAA TGC AGC CAA

Thr Cys Gln Ile Gln Ser Tyr Cys Ala Lys His Leu Lys Cys Ser Gln

1220 1235

AAG TGC GAC CAG AAC AAG TTC AGC GTG AAG TGC TCC TGC TAC GAG GGC
Lys Cys Asp Gln Asn Lys Phe Ser Val Lys Cys Ser Cys Tyr Glu Gly
1240 1245 1250

TGG GTC CTG GAA CCT GAC GGC GAG AGC TGC CGC AGC CTG GAC CCC TTC 4267
Trp Val Leu Glu Pro Asp Gly Glu Ser Cys Arg Ser Leu Asp Pro Phe
1255 1260 1265

AAG CCG TTC ATC ATT TTC TCC AAC CGC CAT GAA ATC CGG CGC ATC GAT

Lys Pro Phe Ile Ile Phe Ser Asn Arg His Glu Ile Arg Arg Ile Asp

1270

1280

CTT CAC AAA GGA GAC TAC AGC GTC CTG GTG CCC GGC CTG CGC AAC ACC

Leu His Lys Gly Asp Tyr Ser Val Leu Val Pro Gly Leu Arg Asn Thr

1285

1290

1295

ATC GCC CTG GAC TTC CAC CTC AGC CAG AGC GCC CTC TAC TGG ACC GAC

Ile Ala Leu Asp Phe His Leu Ser Gln Ser Ala Leu Tyr Trp Thr Asp

1300

1315

GTG GTG GAG GAC AAG ATC TAC CGC GGG AAG CTG CTG GAC AAC GGA GCC

Val Val Glu Asp Lys Ile Tyr Arg Gly Lys Leu Leu Asp Asn Gly Ala

1320

1330

CTG ACT AGT TTC GAG GTG GTG ATT CAG TAT GGC CTG GCC ACA CCC GAG

Leu Thr Ser Phe Glu Val Val Ile Gln Tyr Gly Leu Ala Thr Pro Glu

1335

1340

1345

GGC CTG GCT GTA GAC TGG ATT GCA GGC AAC ATC TAC TGG GTG GAG AGT
Gly Leu Ala Val Asp Trp Ile Ala Gly Asn Ile Tyr Trp Val Glu Ser
1350
1360

AAC CTG GAT CAG ATC GAG GTG GCC AAG CTG GAT GGG ACC CTC CGG ACC
Asn Leu Asp Gln Ile Glu Val Ala Lys Leu Asp Gly Thr Leu Arg Thr
1365
1370
1375

ACC CTG CTG GCC GGT GAC ATT GAG CAC CCA AGG GCA ATC GCA CTG GAT

Thr Leu Leu Ala Gly Asp Ile Glu His Pro Arg Ala Ile Ala Leu Asp

1380

1395

FIG. 14A

(She t 68 of 91)

PIO	Arg	Asp	GIY	11e 1400	CTG Leu	Phe	Trp	Thr	Asp 1405	Trp	Asp	Ala	Ser	Leu 1410	Pro	4699
Arg	116	Glu	Ala 1415	Ala	TCC Ser	Met	Ser :	Gly 1420	Ala	Gly	Arg	Arg	Thr 1425	Val	His	4747
CGG Arg	Glu	ACC Thr 1430	GGC Gly	TCT Ser	GGG Gly	Gly	TGG Trp 1435	Pro	AAC Asn	GGG Gly	Leu	ACC Thr 1440	GTG Val	GAC Asp	TAC Tyr	4795
Leu	Glu 1445	Lys	Arg	Ile	_	Trp 450	Ile	Asp	Ala	Arg	Ser 1455	Asp	Ala	Ile	Tyr	4843
Ser 1460	Ala	Arg	Tyr	Asp	GGC Gly 1465	Ser	Gly	His	Met 1	Glu 1470	Val	Leu	Arg	Gly	His L475	4891
Glu	Phe	Leu	Ser]	His 1480	CCG Pro	Phe	Ala	Val	Thr (485	Leu	Tyr	Gly	Gly	Glu 1490	Val	4939
TAC Tyr	TGG Trp	Thr	GAC Asp 1495	TGG Trp	CGA Arg	ACA Thr	Asn	ACA Thr 1500	CTG Leu	GCT Ala	AAG Lys	Ala	AAC Asn 505	AAG Lys	TGG Trp	4987
ACC Thr	Gly	CAC His US10	AAT Asn	GTC Val	ACC Thr	Val	GTA Val 515	CAG Gln	AGG Arg	ACC Thr	Asn	ACC Thr 520	CAG Gln	CCC Pro	TTT Phe	5035
Asp 1	Leu 525	Gln	Val	Tyr		Pro .530	Ser	Arg	Gln	Pro 1	Met .535	Ala	Pro	Asn	Pro	5083
TGT Cys 1540	GAG Glu	GCC Ala	AAT Asn	Gly	GGC Gly L545	CAG Gln	GCG	CCC Pro	Cys	TCC Ser 1550	CAC His	CTG Leu	TGT Cys	Leu	ATC Ile .555	5131
AAC Asn	TAC Tyr	AAC Asn	Arg	ACC Thr 560	GTG Val	TCC Ser	TGC Cys	Ala	TGC Cys 565	CCC Pro	CAC His	CTC Leu	Met	AAG Lys 1570	CTC Leu	5179
CAC His	AAG Lys	Asp	AAC Asn .575	ACC Thr	ACC	TGC Cys	Tyr	GAG Glu 580	TTT Phe	AAG Lys	AAG Lys	Phe	CTG Leu .585	CTG Leu	TAC Tyr	5227
GCA Ala	Arg	CAG Gln L590	ATG Met	GAG Glu	ATC Ile	Arg	GGT Gly .595	GTG Val	GAC Asp	CTG Leu	Asp	GCT Ala 600	CCC Pro	TAC Tyr	TAC Tyr	5275 -
Asn	TAC Tyr 605	ATC Ile	ATC Ile	TCC Ser	TTC Phe 1	ACG Thr 610	GTG Val	CCC Pro	GAC Asp	Ile	GAC Asp 615	AAC Asn	GTC Val	ACA Thr	GTG Val	5323
CTA Leu 1620	GAC Asp	TAC Tyr	GAT Àsp	Ala	CGC Arg 1625	GAG Glu	CAG Gln	CGT Arg	Val	TAC Tyr 630	TGG Trp	TCT Sér	GAC Asp	Val	CGG Arg 635	5371

FIG. 14A

8449-134 (Sh t 69 of 91)

ACA Thr	CAG Gln	GCC Ala	Ile	AAG Lys 640	CGG Arg	GCC Ala	TTC Phe	Ile	AAC Asn 645	GGC Gly	ACA Thr	GJ Y GGC	Val	GAG Glu .650	ACA Thr	5419
GTC Val	GTC Val	Ser	GCA Ala 655	GAC Asp	TTG Leu	CCA Pro	Asn	GCC Ala 660	CAC His	GGG Gly	CTG Leu	Ala	GTG Val 665	GAC Asp	TGG Trp	5467
	TCC Ser					Trp					Thr					5515
Ile	AAT Asn 1685				Leu					Lys					CAG Gln	5563
	CTG Leu			Pro					Val					Gly		5611
	TAC Tyr		Thr					Ile					Met			5659
	AAT Asn	Arg					Ser					Pro				5707
	ATT Ile					Ser					Ile					5755
His	ACC Thr 1765				Cys					Ser						5803
	GCC Ala			Ser					Ala					Ile		5851
	GAC Asp		Leu					Gln					Met			. 5899
	AGC Ser	Lys		,			Gly					Arg				5947
						Lys					Ser		Gln		GAC Asp	5995 -
His	AAG Lys 1845				Pro		Ser			Asn		Asp			_	6043
	_		-	Thr		Glu			Arg		Cys				GCC Ala 1875	6091

FIG. 14A

(Sheet 70 of 91)

GGC Gly	TAT	AGC Ser	ren	CGG Arg 1880	AGT Ser	GGC Gly	CAG Gln	Gln	GCC Ala 1885	TGC Cys	GAG Glu	GGC Gly	Val	GGT Gly 1890	TCC Ser	6139
Pne	Leu		Tyr 1895	Ser	Val	His	Glu	Gly 1900	Ile	Arg	Gly	Ile	Pro 1905	Leu	Asp	6187
Pro	Asn	1910	Lys	Ser	Asp	Ala	Leu 1915	Val	Pro	Val	Ser]	Gly L920	Thr	Ser	Leu	6235
Ala	vai 1925		TTE	Asp	Pne	H15 1930	Ala	Glu	Asn	Asp :	Thr 1935	Ile	Tyr	Trp	Val	6283
1940	Met	GGC Gly	Leu	Ser :	Thr 1945	Ile	Ser	Arg	Ala	Lys 1950	Arg	Asp	Gln	Thr	Trp 1955	6331
Arg	Glu	GAC Asp	Val	Val 1960	Thr	Asn	Gly	Ile	Gly 1965	Arg	Val	Glu	Gly	Ile 1970	Ala	6379
Val	Asp	,	11e 1975	Ala	Gly	Asn	Ile	Tyr 1980	Trp	Thr	Asp	Gln]	Gly 1985	Phe	Asp	6427
Val	Ile	GAG Glu 1990	Val	Ala	Arg	Leu }	Asn .995	Gly	Ser	Phe	Arg 2	Tyr 2000	Val	Val	Ile	6475
Ser	GIn 2005	GGT Gly	Leu	Asp	Lys 2	Pro 2010	Arg	Ala	Ile	Thr 2	Val 2015	His	Pro	Glu	Lys	6523
Gly 2020	Tyr	TTG Leu	Phe	Trp	Thr 2025	Glu	Tṛp	Gly	Gln 2	Tyr 2030	Pro	Arg	Ile	Glu 2	Arg 2035	6571
Ser	Arg	CTA Leu	Asp 2	Gly 2040	Thr	Glu	Arg	Val 2	Val 2045	Leu	Val	Asn	Val	Ser 2050	Ile	6619
Ser	Trp		Asn 2055	Gly	Ile	Ser	Val 2	Asp 2060	Tyr	Gln	Asp	Gly 2	Lys 2065	Leu	Tyr	6667
TGG Trp	Cys	GAT Asp 2070	GCA Ala	CGG Arg	ACA Thr	Asp	AAG Lys 075	ATT Ile	GAA Glu	CGG Arg	Ile	GAC Asp 1080	CTG Leu	GAG Glu	ACA Thr	6715 -
GLY	GAG Glu 2085	AAC Asn	CGC Arg	GAG Glu	Val	GTT Val 090	CTG Leu	TCC Ser	AGC Ser	Asn	AAC Asn 2095	ATG Met	GAC Aşp	ATG Met	TTT Phe	67 _, 63
TCA Ser 2100	GTG Val	TCT Ser	GTG Val	Phe	GAG Glu 2105	GAT Asp	TTC Phe	ATC Ile	Tyr	TGG Trp 2110	AGT Ser	GAC Asp	AGG Arg	Thr	CAT His 2115	6811

FIG. 14A

(She t 71 of 91)

GCC Ala	AAC Asn	GGC Gly	Ser	ATC Ile 2120	AAG Lys	CGC Arg	GGG Gly	Ser	AAA Lys 2125	GAC Asp	AAT Asn	GCC Ala	Thr	GAC Asp 2130	TCC Ser	6859
GTG Val	CCC Pro	Leu	CGA Arg 2135	ACC Thr	GGC Gl _y	ATC Ile	Gly	GTC Val 2140	CAG Gln	CTT Leu	AAA Lys	Asp	ATC Ile 2145	AAA Lys	GTC Val	6907
TTC Phe	AAC Asn	CGG Arg 2150	GAC Asp	CGG Arg	CAG Gln	Lys	GGC Gly 2155	ACC Thr	AAC Asn	GTG Val	Cys	GCG Ala 2160	GTG Val	GCC Ala	AAT Asn	6955
Gly	GGG Gly 2165	TGC Cys	CAG Gln	CAG Gln	Leu	TGC Cys 2170	CTG Leu	TAC Tyr	CGG Arg	Gly	CGT Arg 2175	GGG Gly	CAG Gln	CGG Arg	GCC Ala	7003·
TGC Cys 2180	GCC Ala	TGT Cys	GCC Ala	His	GGG Gly 2185	ATG Met	CTG Leu	GCT Ala	Glu	GAC Asp 2190	GGA Gly	GCA Ala	TCG Ser	Cys	CGC Arg 2195	7051
GAG Glu	TAT Tyr	GCC	Gly	TAC Tyr 2200	CTG Leu	CTC Leu	TAC Tyr	Ser	GAG Glu 205	CGC Arg	ACC Thr	ATT Ile	Leu	AAG Lys 2210	AGT Ser	7099
ATC Ile	CAC His	Leu	TCG Ser 2215	GAT Asp	GAG Glu	CGC Arg	Asn	CTC Leu 2220	AAT Asn	GCG Ala	CCC Pro	Val	CAG Gln 2225	CCC Pro	TTC Phe	7147
GAG Gl u	GAC Asp	CCT Pro 230	GAG Glu	CAC His	ATG Met	Lys	AAC Asn 235	GTC Val	ATC Ile	GCC Ala	Leu	GCC Ala 240	TTT Phe	GAC Asp	TAC Tyr	7195
Arg	GCA Ala 2245	GGC Gly	ACC Thr	TCT Ser	Pro	GGC Gly 2250	ACC Thr	CCC Pro	AAT Asn	Arg	ATC 11e 255	TTC Phe	TTC Phe	AGC Ser	GAC Asp	7243
ATC 11e 2260	CAC His	TTT Phe	GGG Gly	Asn	ATC Ile 2265	CAA Gln	CAG Gln	ATC Ile	Asn	GAC Asp 270	GAT Asp	GGC Gly	TCC Ser	Arg	AGG Arg 275	7291
ATC Ile	ACC Thr	ATT Ile	Val	GAA Glu 280	AAC Asn	GTG Val	Gly GGC	Ser	GTG Val 285	GAA Glu	GGC Gly	CTG Leu	Ala	TAT Tyr 2290	CAC His	.7339 ·
CGT Arg	GGC Gly	Trp	GAC Asp 295	ACT Thr	CTC Leu	TAT Tyr	Trp	ACA Thr 300	AGC Ser	TAC Tyr	ACG Thr	Thr	TCC Ser	ACC Thr	ATC Ile	7387
ACG Thr	CGC Arg 2	CAC His 310	ACA Thr	GTG Val	GAC Asp	Gln	ACC Thr 2315	CGC Arg	CCA Pro	GGG Gly	Ala	TTC Phe	GAG Glu	CGT Arg	GAG Glu	7435
Thr	GTC Val 2325	ATC Ile	ACT Thr	ATG Met	Ser	GGA Gly 2330	GAT Asp	GAC Asp	CAC His	Pro	CGG Arg	GCC Ala	TTC Phe	GTT Val	TTG Leu	7483
GAC Asp 2340	GAG Glu	TGC Cys	CAG Gln	Asn	CTC Leu 2345	ATG Met	TTC Phe	TGG Trp	Thr	AAC Asn 2350	TGG Trp	AAT Asn	GAG Glu	Gln	CAT His 2355	7531



(Sh et 72 of 91)

CCC Pro	AGC Ser	ATC Ile	Met	CGG Arg 2360	GCG Ala	GCG Ala	CTC Leu	Ser	GGA Gly 2365	GCC Ala	AAT Asn	GTC Val	Leu	ACC Thr 2370	CTT Leu	7579
ATC Ile	GAG Glu	Lys	GAC Asp 2375	ATC Ile	CGT Arg	ACC Thr	Pro	AAT Asn 0862	GGC Gly	CTG Leu	GCC Ala	Ile	GAC Asp 2385	CAC His	CGT Arg	7627
GCC Ala	Glu	AAG Lys 2390	CTC Leu	TAC Tyr	TTC Phe	Ser	GAC Asp 2395	GCC Ala	ACC Thr	CTG Leu	Asp	AAG Lys 2400	ATC Ile	GAG Glu	CGG Arg	7675
Cys	GAG Glu 2405	TAT Tyr	GAC Asp	GGC Gly	TCC Ser	CAC His 2410	C GC A rg	TAT Tyr	GTG Val	Ile	CTA Leu 2415	AAG Lys	TCA Ser	GAG Glu	CCT Pro	7723
GTC Val 2420	CAC His	CCC Pro	TTC Phe	Gly	CTG Leu 2425	GCC Ala	GTG Val	TAT Tyr	Gly	GAG Glu 430	CAC His	ATT Ile	TTC Phe	Trp	ACT Thr 2435	7771
GAC Asp	TGG Trp	GTG Val	Arg	CGG Arg 2440	GCA Ala	GTG Val	CAG Gln	Arg	GCC Ala 2445	AAC Asn	AAG Lys	CAC His	Val	GGC Gly 2450	AGC Ser	7819
AAC Asn	ATG Met	Lys	CTG Leu 2455	CTG Leu	CGC Arg	GTG Val	Asp	ATC Ile 2460	CCC Pro	CAG Gln	CAG Gln	Pro	ATG Met 465	GJ Y	ATC Ile	7867
	Ala				GAC Asp	Thr					Leu					7915
Ile					TGC Cys 2					Leu						7963
				Ser	TGC Cys 2505				Arg					Asp		8011
			Ala		TAA Asn			Cys					Glu			8059
		Asn			TGC Cy s		Asn					Cys			GTC Val	8107
	His				AAG Lys	Ser					Ser					8155
Arg					ACT Thr					Ser						8203
				Trp	TGC Cys 2585				Asp					Gly		8251 .

FIG. 14A

8449-134 (Sheet 73 of 91)

0-	TTU	- 10	•										1	•••		• • • • • • • • • • • • • • • • • • • •
	GAG Glu		Pro					Ala					Glu			8299
	CGG Arg	Asp		Thr			Gly					Cys				8347
GTG Val	GAT Asp	TGT Cys 630	GAG Glu	GAC Asp	GCC Ala	Ser	GAT Asp 2635	GAG Glu	ATG Met	AAC Asn	Cys	AGT Ser 2640	GCC Ala	ACC Thr	GAC Asp	8395
Cys	AGC Ser 2645	AGC Ser	TAC Tyr	TTC Phe	Arg	CTG Leu 2650	GGC Gly	GTG Val	AAG Lys	Gly	GTG Val 2655	CTC Leu	TTC Phe	CAG Gln	CCC Pro	8443
	GAG Glu			Ser					Pro					Asp		8491
GCC Ala	TAA neA	GAC Asp	Cys	GGG Gly 2680	GAC Asp	TAC Tyr	AGT Ser	Asp	GAG Glu 685	CGC Arg	GAC Asp	TGC Cys	Pro	GGT Gly 2690	Val	8539
AAA Lys	CGC Arg	Pro	AGA Arg 2695	TGC Cys	CCT Pro	CTG Leu	Asn	TAC Tyr 2700	TTC Phe	GCC Ala	TGC Cys	Pro	AGT Ser 2705	G1 y	CGC Arg	8587
TGC Cys	Ile	Pro	ATG Met	Ser	Trp	Thr	Cys	Asp	AAA Lys	GAG Glu	Asp	GAC Asp 2720	Cys	GAA Glu	CAT His	8635
Gly	GAG Glu 2725	GAC Asp	GAG Glu	ACC Thr	His	TGC Cys 2730	AAC Asn	AAG Lys	TTC Phe	Cys	TCA Ser 2735	GAG Glu	GCC Ala	CAG Gln	TTT Phe	8683
GAG Glu 2740	Cys	CAG Gln	AAC Asn	His	CGC Arg 2745	Cys	ATC Ile	TCC Ser	Lys	CAG Gln 2750	Trp	CTG Leu	TGT Cys	GAC Asp	GGC Gly 2755	8731
AGC Ser	GAT Asp	GAC Asp	Cys	GGG Gly 2760	Asp	GGC Gly	TCA Ser	Asp	GAG Glu 2765	Ala	GCT Ala	CAC	TGT Cys	GA! Glu 2770	GGC Gly	8779
AAG Lys	ACG Thr	TGC Cys	GGC Gly 2775	Pro	TCC	TCC Ser	Phe	TCC Ser 2780	Cys	CCT Pro	GGC Gly	ACC Thr	CAC His 2785	Va.	TGC L Cys	8827
GT(Val	CCC	GAG Glu 2790	. Arg	TGG Trp	CTC Lev	TGI Cys	GAC Asp 2795	o Gly	GAC Asp	AAA Lys	A GAC	TG1 Cys 2800	s Ala	GA' A Asj	T GGT p Gly	8875 ·
GC# Ala	A GAC A Asp 2805	Glu	AGC Ser	ATC	GCA Ala	A GCT A Ala 2810	a Gly	TGC Cys	TTG Leu	TAC Tyr	AAC Asr 2815	Se	C ACT	T TG	T GAC s Asp	8923
GA(As ₁ 282	Arç	GAC Glu	TTC	ATC Met	TG0 Cys 282	s Gl	AAG A Ası	C CGC	CAC Glr	TG0 Cys 2830	s Ile	C CC	C AAG o Ly:	G CA s Hi	C TTC s Phe 2835	8971

8449-134 GTG TGT GAC CAC GAC CGT GAC TGT GCA GAT GGC TCT GAT GAG TCC CCC 9019 Val Cys Asp His Asp Arg Asp Cys Ala Asp Gly Ser Asp Glu Ser Pro 2840 CAG TGT GAG TAC CCG ACC TGC GGC CCC AGT GAG TTC CGC TGT GCC AAT 9067

GGG CGC TGT CTG AGC TCC CGC CAG TGG GAG TGT GAT GGC GAG AAT GAC

Gly Arg Cys Leu Ser Ser Arg Gln Trp Glu Cys Asp Gly Glu Asn Asp

2870

2880

Glu Cys Glu Tyr Pro Thr Cys Gly Pro Ser Glu Phe Arg Cys Ala Asn

2860

2855

TGC CAC GAC CAG AGT GAC GAG GCT CCC AAG AAC CCA CAC TGC ACC AGC

9163
Cys His Asp Gln Ser Asp Glu Ala Pro Lys Asn Pro His Cys Thr Ser

2885

2890

2895

CCA GAG CAC AAG TGC AAT GCC TCG TCA CAG TTC CTG TGC AGC AGT GGG 9211
Pro Glu His Lys Cys Asn Ala Ser Ser Gln Phe Leu Cys Ser Ser Gly
2900 2915

CGC TGT GTG GCT GAG GCA CTG CTC TGC AAC GGC CAG GAT GAC TGT GGC 9259
Arg Cys Val Ala Glu Ala Leu Leu Cys Asn Gly Gln Asp Asp Cys Gly
2920 2925 2930

GAC AGC TCG GAC GAG CGT GGC TGC CAC ATC AAT GAG TGT CTC AGC CGC 9307
Asp Ser Ser Asp Glu Arg Gly Cys His Ile Asn Glu Cys Leu Ser Arg
2935 2940 2945

AAG CTC AGT GGC TGC AGC CAG GAC TGT GAG GAC CTC AAG ATC GGC TTC 9355

Lys Leu Ser Gly Cys Ser Gln Asp Cys Glu Asp Leu Lys Ile Gly Phe 2950 2960

AAG TGC CGC TGT CGC CCT GGC TTC CGG CTG AAG GAT GAC GGC CGG ACG 9403 Lys Cys Arg Cys Arg Pro Gly Phe Arg Leu Lys Asp Asp Gly Arg Thr 2965 2970 2975

TGT GCT GAT GTG GAC GAG TGC AGC ACC ACC TTC CCC TGC AGC CAG CGC 9451 .

Cys Ala Asp Val Asp Glu Cys Ser Thr Thr Phe Pro Cys Ser Gln Arg
2980 2995 .

TGC ATC AAC ACC CAT GGC AGC TAT AAG TGT CTG TGT GTG GAG GGC TAT 9499
Cys Ile Asn Thr His Gly Ser Tyr Lys Cys Leu Cys Val Glu Gly Tyr .
3000 3005 3010

GCA CCC CGC GGC GGC GAC CCC CAC AGC TGC AAG GCT GTG ACT GAC GAG
Ala Pro Arg Gly Gly Asp Pro His Ser Cys Lys Ala Val Thr Asp Glu
3015 3020 3025

GAA CCG TTT CTG ATC TTC GCC AAC CGG TAC TAC CTG CGC AAG CTC AAC

9595
Glu Pro Phe Leu Ile Phe Ala Asn Arg Tyr Tyr Leu Arg Lys Leu Asn
3030

3040

CTG GAC GGG TCC AAC TAC ACG TTA CTT AAG CAG GGC CTG AAC AAC GCC
Leu Asp Gly Ser Asn Tyr Thr Leu Leu Lys Gln Gly Leu Asn Asn Ala
3045
3050
3055

GTT GCC TTG GAT TTT GAC TAC CGA GAG CAG ATG ATC TAC TGG ACA GAT 9691

Val Ala Leu Asp Phe Asp Tyr Arg Glu Gln Met Ile Tyr Trp Thr Asp

3060 3075

FIG. 14A

(She t 75 of 91)

GTG ACC ACC CAG GGC AGC ATG ATC CGA AGG ATG CAC CTT AAC GGG AGC Val Thr Thr Gln Gly Ser Met Ile Arg Arg Met His Leu Asn Gly Ser 3080 3085 3090	9739
AAT GTG CAG GTC CTA CAC CGT ACA GGC CTC AGC AAC CCC GAT GGG CTG Asn Val Gln Val Leu His Arg Thr Gly Leu Ser Asn Pro Asp Gly Leu 3095 3100 3105	9787
GCT GTG GAC TGG GTG GGT GGC AAC CTG TAC TGG TGC GAC AAA GGC CGG Ala Val Asp Trp Val Gly Gly Asn Leu Tyr Trp Cys Asp Lys Gly Arg 3110 3115 3120	9835
GAC ACC ATC GAG GTG TCC AAG CTC AAT GGG GCC TAT CGG ACG GTG CTG Asp Thr Ile Glu Val Ser Lys Leu Asn Gly Ala Tyr Arg Thr Val Leu 3125 3130 3135	9883
GTC AGC TCT GGC CTC CGT GAG CCC AGG GCT CTG GTG GAT GTG CAG Val Ser Ser Gly Leu Arg Glu Pro Arg Ala Leu Val Val Asp Val Gln 3140 3155	9931
AAT GGG TAC CTG TAC TGG ACA GAC TGG GGT GAC CAT TCA CTG ATC GGC Asn Gly Tyr Leu Tyr Trp Thr Asp Trp Gly Asp His Ser Leu Ile Gly 3165 3170	9979
CGC ATC GGC ATG GAT GGG TCC AGC CGC AGC GTC ATC GTG GAC ACC AAG Arg Ile Gly Met Asp Gly Ser Ser Arg Ser Val Ile Val Asp Thr Lys 3175 3180 3185	10027
ATC ACA TGG CCC AAT GGC CTG ACG CTG GAC TAT GTC ACT GAG CGC ATC Ile Thr Trp Pro Asn Gly Leu Thr Leu Asp Tyr Val Thr Glu Arg Ile 3190 3200	10075
TAC TGG GCC GAC GCC CGC GAG GAC TAC ATT GAA TTT GCC AGC CTG GAT Tyr Trp Ala Asp Ala Arg Glu Asp Tyr Ile Glu Phe Ala Ser Leu Asp 3205 3210 3215	10123
GGC TCC AAT CGC CAC GTT GTG CTG AGC CAG GAC ATC CCG CAC ATC TTT Gly Ser Asn Arg His Val Val Leu Ser Gln Asp Ile Pro His Ile Phe 3220 3235	10171
GCA CTG ACC CTG TTT GAG GAC TAC GTC TAC TGG ACC GAC TGG GAA ACA Ala Leu Thr Leu Phe Glu Asp Tyr Val Tyr Trp Thr Asp Trp Glu Thr 3240 3245 3250	10219
AAG TCC ATT AAC CGA GCC CAC AAG ACC ACG GGC ACC AAC AA	10267
CTC ATC AGC ACG CTG CAC CGG CCC ATG GAC CTG CAT GTC TTC CAT GCC Leu Ile Ser Thr Leu His Arg Pro Met Asp Leu His Val Phe His Ala 3270 3280	10315
CTG CGC CAG CCA GAC GTG CCC AAT CAC CCC TGC AAG GTC AAC AAT GGT Leu Arg Gln Pro Asp Val Pro Asn His Pro Cys Lys Val Asn Asn Gly 3285 3290 3295	10363
•	

8449-134 (Sh t 76 of 91) GCC TGC CCC ACC AAC TTC TAC CTG GGC AGC GAT GGG CGC ACC TGT GTG Ala Cys Pro Thr Asn Phe Tyr Leu Gly Ser Asp Gly Arg Thr Cys Val 3320 TCC AAC TGC ACG GCT AGC CAG TTT GTA TGC AAG AAC GAC AAG TGC ATC 10507 Ser Asn Cys Thr Ala Ser Gln Phe Val Cys Lys Asn Asp Lys Cys Ile 3335 3340 CCC TTC TGG TGG AAG TGT GAC ACC GAG GAC GAC TGC GGG GAC CAC TCA 10555 Pro Phe Trp Trp Lys Cys Asp Thr Glu Asp Asp Cys Gly Asp His Ser 3350 3355 GAC GAG CCC CCG GAC TGC CCT GAG TTC AAG TGC CGG CCC GGA CAG TTC 10603 Asp Glu Pro Pro Asp Cys Pro Glu Phe Lys Cys Arg Pro Gly Gln Phe 3365 CAG TGC TCC ACA GGT ATC TGC ACA AAC CCT GCC TTC ATC TGC GAT GGC 10651 Gln Cys Ser Thr Gly Ile Cys Thr Asn Pro Ala Phe Ile Cys Asp Gly 3385 3390 GAC AAT GAC TGC CAG GAC AAC AGT GAC GAG GCC AAC TGT GAC ATC CAC 10699 Asp Asn Asp Cys Gln Asp Asn Ser Asp Glu Ala Asn Cys Asp Ile His -3400 3405 3410 GTC TGC TTG CCC AGT CAG TTC AAA TGC ACC AAC ACC AAC CGC TGT ATT 10747 Val Cys Leu Pro Ser Gln Phe Lys Cys Thr Asn Thr Asn Arg Cys Ile 3415 3420 3425 CCC GGC ATC TTC CGC TGC AAT GGG CAG GAC AAC TGC GGA GAT GGG GAG 10795 Pro Gly Ile Phe Arg Cys Asn Gly Gln Asp Asn Cys Gly Asp Gly Glu 3430 GAT GAG AGG GAC TGC CCC GAG GTG ACC TGC GCC CCC AAC CAG TTC CAG 10843 Asp Glu Arg Asp Cys Pro Glu Val Thr Cys Ala Pro Asn Gln Phe Gln 3445 3450 TGC TCC ATT ACC AAA CGG TGC ATC CCC CGG GTC TGG GTC TGC GAC CGG 10891 . Cys Ser Ile Thr Lys Arg Cys Ile Pro Arg Val Trp Val Cys Asp Arg 3460 3465 3470 GAC AAT GAC TGT GTG GAT GGC AGT GAT GAG CCC GCC AAC TGC ACC CAG 10939 Asp Asn Asp Cys Val Asp Gly Ser Asp Glu Pro Ala Asn Cys Thr Gln 3480 3485 3490 ATG ACC TGT GGT GTG GAC GAG TTC CGC TGC AAG GAT TCG GGC CGC TGC 10987 Met Thr Cys Gly Val Asp Glu Phe Arg Cys Lys Asp Ser Gly Arg Cys 3495 ATC CCA GCG CGT TGG AAG TGT GAC GGA GAG GAT GAC TGT GGG GAT GGC 11035 Ile Pro Ala Arg Trp Lys Cys Asp Gly Glu Asp Asp Cys Gly Asp Gly 3510 3515 TCG GAT GAG CCC AAG GAA GAG TGT GAT GAA CGC ACC TGT GAG CCA TAC 11083 Ser Asp Glu Pro Lys Glu Glu Cys Asp Glu Arg Thr Cys Glu Pro Tyr 3525 3530 3535 11131 CAG TTC CGC TGC AAG AAC AAC CGC TGC GTG CCC GGC CGC TGG CAG TGC Gln Phe Arg Cys Lys Asn Asn Arg Cys Val Pro Gly Arg Trp Gln Cys

FIG. 14A

3550

3545

8449-134 (Sh et 77 of 91) GAC TAC GAC AAC GAT TGC GGT GAC AAC TCC GAT GAA GAG AGC TGC ACC Asp Tyr Asp Asn Asp Cys Gly Asp Asn Ser Asp Glu Glu Ser Cys Thr 3560 CCT CGG CCC TGC TCC GAG AGT GAG TTC TCC TGT GCC AAC GGC CGC TGC 11227 Pro Arg Pro Cys Ser Glu Ser Glu Phe Ser Cys Ala Asn Gly Arg Cys 3575 3580 ATC GCG GGG CGC TGG AAA TGC GAT GGA GAC CAC GAC TGC GCG GAC GGC 11275 Ile Ala Gly Arg Trp Lys Cys Asp Gly Asp His Asp Cys Ala Asp Gly 3595 3590 3600 TCG GAC GAG AAA GAC TGC ACC CCC CGC TGT GAC ATG GAC CAG TTC CAG 11323 Ser Asp Glu Lys Asp Cys Thr Pro Arg Cys Asp Met Asp Gln Phe Gln 3605 3610 TGC AAG AGC GGC CAC TGC ATC CCC CTG CGC TGG CGC TGT GAC GCA GAC 11371 Cys Lys Ser Gly His Cys Ile Pro Leu Arg Trp Arg Cys Asp Ala Asp 3620 3625 3630 GCC GAC TGC ATG GAC GGC AGC GAC GAG GCC TGC GGC ACT GGC GTG 11419 Ala Asp Cys Met Asp Gly Ser Asp Glu Glu Ala Cys Gly Thr Gly Val

3640 3645 3650 CGG ACC TGC CCC CTG GAC GAG TTC CAG TGC AAC AAC ACC TTG TGC AAG 11467 Arg Thr Cys Pro Leu Asp Glu Phe Gln Cys Asn Asn Thr Leu Cys Lys 3655 3660 CCG CTG GCC TGG AAG TGC GAT GGC GAG GAT GAC TGT GGG GAC AAC TCA 11515 Pro Leu Ala Trp Lys Cys Asp Gly Glu Asp Asp Cys Gly Asp Asn Ser 3675 3670 3680 GAT GAG AAC CCC GAG GAG TGT GCC CGG TTC GTG TGC CCT CCC AAC CGG 11563 Asp Glu Asn Pro Glu Glu Cys Ala Arg Phe Val Cys Pro Pro Asn Arg 3685 3690 CCC TTC CGT TGC AAG AAT GAC CGC GTC TGT CTG TGG ATC GGG CGC CAA 11611 . Pro Phe Arg Cys Lys Asn Asp Arg Val Cys Leu Trp Ile Gly Arg Gln 3700 3705 3710 3715 11659 TGC GAT GGC ACG GAC AAC TGT GGG GAT GGG ACT GAT GAA GAG GAC TGT Cys Asp Gly Thr Asp Asn Cys Gly Asp Gly Thr Asp Glu Glu Asp Cys 3720 3725 3730 GAG CCC CCC ACA GCC CAC ACC CAC TGC AAA GAC AAG AAG GAG TTT 11707 Glu Pro Pro Thr Ala His Thr Thr His Cys Lys Asp Lys Lys Glu Phe 3735 3740 3745 11755 CTG TGC CGG AAC CAG CGC TGC CTC TCC TCC CTG CGC TGC AAC ATG Leu Cys Arg Asn Gln Arg Cys Leu Ser Ser Ser Leu Arg Cys Asn Met 3755 3750 3760 11803 TTC GAT GAC TGC GGG GAC GGC TCT GAC GAG GAG GAC TGC AGC ATC GAC Phe Asp Asp Cys Gly Asp Gly Ser Asp Glu Glu Asp Cys Ser Ile Asp 3770 3765 3775 CCC AAG CTG ACC AGC TGC GCC ACC AAT GCC AGC ATC TGT GGG GAC GAG 11851 Pro Lys Leu Thr Ser Cys Ala Thr Asn Ala Ser Ile Cys Gly Asp Glu

FIG. 14A

3790

3785

8449-134 (Sheet 78 of 91)

GCA CGC TGC GTG CGC ACC GAG AAA GCG GCC TAC TGT GCC TGC CGC TCG Ala Arg Cys Val Arg Thr Glu Lys Ala Ala Tyr Cys Ala Cys Arg Ser 3800	11899
GGC TTC CAC ACC GTG CCC GGC CAG CCC GGA TGC CAA GAC ATC AAC GAG Gly Phe His Thr Val Pro Gly Gln Pro Gly Cys Gln Asp Ile Asn Glu 3815	11947
TGC CTG CGC TTC GGC ACC TGC TCC CAG CTC TGC AAC AAC ACC AAG GGC Cys Leu Arg Phe Gly Thr Cys Ser Gln Leu Cys Asn Asn Thr Lys Gly 3835	11995
GGC CAC CTC TGC AGC TGC GCT CGG AAC TTC ATG AAG ACG CAC AAC ACC Gly His Leu Cys Ser Cys Ala Arg Asn Phe Met Lys Thr His Asn Thr 3845 3850 3855	12043
TGC AAG GCC GAA GGC TCT GAG TAC CAG GTC CTG TAC ATC GCT GAT GAC Cys Lys Ala Glu Gly Ser Glu Tyr Gln Val Leu Tyr Ile Ala Asp Asp 3860 3875	12091
AAT GAG ATC CGC AGC CTG TTC CCC GGC CAC CCC CAT TCG GCT TAC GAG Asn Glu Ile Arg Ser Leu Phe Pro Gly His Pro His Ser Ala Tyr Glu 3880 3885 3890	12139
CAG GCA TTC CAG GGT GAC GAG AGT GTC CGC ATT GAT GCT ATG GAT GTC Gln Ala Phe Gln Gly Asp Glu Ser Val Arg Ile Asp Ala Met Asp Val 3895 3900 3905	12187
CAT GTC AAG GCT GGC CGT GTC TAT TGG ACC AAC TGG CAC ACG GGC ACC His Val Lys Ala Gly Arg Val Tyr Trp Thr Asn Trp His Thr Gly Thr 3910 3920	12235
ATC TCC TAC CGC AGC CTG CCA CCT GCT GCG CCT CCT ACC ACT TCC AAC Ile Ser Tyr Arg Ser Leu Pro Pro Ala Ala Pro Pro Thr Thr Ser Asn 3925 3935	12283
CGC CAC CGG CGA CAG ATT GAC CGG GGT GTC ACC CAC CTC AAC ATT TCA Arg His Arg Arg Gln Ile Asp Arg Gly Val Thr His Leu Asn Ile Ser 3940 3950 3955	12331
GGG CTG AAG ATG CCC AGA GGC ATC GCC ATC GAC TGG GTG GCC GGA AAC Gly Leu Lys Met Pro Arg Gly Ile Ala Ile Asp Trp Val Ala Gly Asn 3960 3970	12379
GTG TAC TGG ACC GAC TCG GGC CGA GAT GTG ATT GAG GTG GCG CAG ATG Val Tyr Trp Thr Asp Ser Gly Arg Asp Val Ile Glu Val Ala Gln Met 3975	12427
AAG GGC GAG AAC CGC AAG ACG CTC ATC TCG GGC ATG ATT GAC GAG CCC Lys Gly Glu Asn Arg Lys Thr Leu Ile Ser Gly Met Ile Asp Glu Pro 3990 3995 4000	12475
CAC GCC ATT GTG GTG GAC CCA CTG AGG GGG ACC ATG TAC TGG TCA GAC His Ala Ile Val Val Asp Pro Leu Arg Gly Thr Met Tyr Trp Ser Asp 4005 4015	12523
TGG GGC AAC CAC CCC AAG ATT GAG ACG GCA GCG ATG GAT GGG ACG CTT Trp Gly Asn His Pro Lys Ile Glu Thr Ala Ala Met Asp Gly Thr Leu 4020 4035	12571

FIG. 14A

8449-134 (She t 79 of 91)

*** CGG Arg	GAG Glu	ACA Thr	ьeu	GTG Val 4040	GIH	GAC Asp	AAC Asn	Ile	CAG Gln 4045	TGG Trp	CCC Pro	ACA Thr	Gly	CTG Leu 4050	GCC Ala	12619
Val	Asp	TYF.	4055	ASN	. GIU	Arg	Leu	Tyr 4060	Trp	Ala	Asp	Ala	Lys 4065	Leu		12667
AGI	rre	4070	ser	ire	CGG Arg	Leu	Asn 4075	Gly	Thr	Asp	Pro	11e 4080	Val	Ala	Ala	12715
Asp	3er 4085	Lys	Arg	GIA		Ser 1090	His	Pro	Phe	Ser	11e 4095	Asp	Val	Phe	Glu	12763
4100	Tyr	116	Tyr	GTA	GTC Val 4105	Thr	Tyr	Ile	Asn	Asn 1110	Arg	Val	Phe	Lys	Ile 4115	12811
nis	ьys	Pne	GIÀ	H1S 120	AGC Ser	Pro	Leu	Val	Asn 125	Leu	Thr	Gly	Gly	Leu 4130	Ser	12859
CAC His	GCC Ala	Ser	GAC Asp 1135	GTG Val	GTC Val	CTT Leu	Tyr	CAT His 1140	CAG Gln	CAC His	AAG Lys	Gln	CCC Pro	GAA Glu	GTG Val	12907
ACC Thr	Asn	CCA Pro 1150	TGT Cys	GAC Asp	CGC Arg	Lys	AAA Lys 1155	TGC Cys	GAG Glu	TGG Trp	Leu	TGC Cys 1160	CTG Leu	CTG Leu	AGC Ser	12955
Pro	AGT Ser 1165	GGG Gly	CCT Pro	GTC Val	TGC Cys 4	ACC Thr 170	TGT Cys	CCC Pro	AAT Asn	Gly	AAG Lys 1175	CGG Arg	CTG Leu	GAC Asp	AAC Asn	13003
4180	Inr	Cys	val	Pro 4	GTG Val 1185	Pro	Ser	Pro	Thr	Pro 190	Pro	Pro	Asp	Ala 4	Pro 1195	13051 •
CGG Arg	CCT Pro	GGA Gly	Thr	TGT Cys 200	AAC Asn	CTG Leu	CAG Gln	Cys	TTC Phe 205	AAC Asn	GGT Gly	GJ A CCC	Ser	TGT Cys 1210	TTC Phe	13099
CTC Leu	AAT Asn	Ala	CGG Arg 215	AGG Arg	CAG Gln	CCC Pro	Lys	TGC Cys 220	CGC Arg	TGC Cys	CAA Gln	Pro	CGC Arg 1225	TAC Tyr	ACG Thr	13147
GGT Gly	Asp	AAG Lys 230	TGT Cys	GAA Glu	CTG Leu	Asp	CAG Gln 235	TGC Cys	TGG Trp	GAG Glu	His	TGT Cys 1240	CGC Arg	TAA Asn	GLY GLY	13195 -
CTA	ACC Thr 1245	TGT Cys	GCT Ala	GCC Ala	TCC Ser	CCC Pro 250	TCT Ser	GGC Gly	ATG Met	Pro	ACG Thr 255	TGC Cys	CGG Arg	TGC Cys	CCC Pro	13243
ACG Thr 4260	GGC Gly	TTC Phe	ACG Thr	GIÀ	CCC Pro 265	AAA Lys	TGC Cys	ACC Thr	Gln	CAG Gln 270	GTG Val	TGT Cys	GĆG Ala	Gly	TAC Tyr 1275	13291

FIG. 14A

(Sheet 80 of 91)

0,1	, ,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,	AAC Asn	ASII	4280	Int	Cys ,	inr	Val	Asn 4285	Gln	Gly	Asn	Gln	Pro 4290	G1n	13339
Cys	, Arg		4295	ĖĽO	GIA	Pne	reu	Gly 4300	Asp	Arg	Cys	Gln	Tyr 4305	Arg	Gln	13387
Cys	Ser	GGC Gly 4310	ıyı	Cys	GIU	Asn	4315	GIA	Thr	Cys	Gln	Met 4320	Ala	Ala	Asp	13435
Gly	4325		GIN	Cys	Arg	Cys 1330	Thr	Ala	Tyr	Phe	Glu 4335	Gly	Ser	Arg	Cys	13483
4340	val	AAC Asn	Lys	Cys 4	Ser 1345	Arg	Cys	Leu	Glu	Gly 4350	Ala	Cys	Val	Val	Asn 1355	13531
БУЗ	GIN	AGT Ser	4	360	val	Thr	Cys	Asn	Cys 1365	Thr	Asp	Gly	Arg	Val 4370	Ala	13579
PIO	Sei		1375	rnr	Cys	Val	GIY 4	His 1380	Cys	Ser	Asn	Gly 4	Gly 385	Ser	Cys	13627
inr	Met	AAC Asn 4390	ser	ьуs	Met	Met 4	9ro 395	Glu	Cys	Gln	Cys 4	Pro 400	Pro	His	Met	13675
IIIL	4405	CCC Pro	Arg	Cys	4	410	His	Val	Phe	Ser 4	Gln 415	Gln	Gln	Pro	Gly	13723
4420	rre	GCC Ala	ser	11e 4	125	He	Pro	Leu	Leu 4	Leu 430	Leu	Leu	Leu	Leu 4	Val 435	13771
Leu	vai	GCC Ala	G1 y 4	440	Val	Phe	Trp	Tyr 4	Lys 445	Arg	Arg	Val	Gln 4	Gly 450	Ala	13819
ьys	GIA		455	HIS	GIn	Arg	Met 4	Thr 460	Asn _.	Gly	Ala	Met 4	Asn 465	Val	Glu	13867
ATT Ile	етА	AAC Asn 470	CCC . Pro	ACC Thr	TAC Tyr	Lys	ATG Met 475	TAC Tyr	GAA Glu	GGC Gly	Gly	GAG Glu 480	CCT Pro	GAT Asp	GAT Asp	13915
val	GGA Gly 1485	GGC Gly	CTA Leu	CTG Leu	Asp	GCT Ala 490	GAC Asp	TTT Phe	GCC Ala	Leu	GAC Asp 495	CCT Pro	GAC Asp	AAG Lys	CCC Pro	13963
ACC Thr 4500	AAC Asn	TTC .	ACC . Thr	Asn	CCC Pro 505	GTG Val	TAT Tyr	GCC Ala	Thr	CTC Leu 510	TAC Tyr	ATG Met	GGG Gly	Gly	CAT His 515	14011

FIG. 14A

14896

AAAAAA

GGC AGT CGC CAC TCC CTG GCC AGC ACG GAC GAG AAG CGA GAA CTC CTG 14059 Gly Ser Arg His Ser Leu Ala Ser Thr Asp Glu Lys Arg Glu Leu Leu 4520 4525 GGC CGG GGC CCT GAG GAC GAG ATA GGG GAC CCC TTG GCA TAGGGCCCTG CC 14110 CCGTCGGACT GCCCCCAGAA AGCCTCCTGC CCCCTGCCGG TGAAGTCCTT CAGTGAGCCC Gly Arg Gly Pro Glu Asp Glu Ile Gly Asp Pro Leu Ala CTCCCCAGCC AGCCCTTCCC TGGCCCCGCC GGATGTATAA ATGTAAAAAT GAAGGAATTA CATTTTATAT GTGAGCGAGC AAGCCGGCAA GCGAGCACAG TATTATTTCT CCATCCCTC CCTGCCTGCT CCTTGGCACC CCCATGCTGC CTTCAGGGAG ACAGGCAGGG AGGGCTTGGG 14350 GCTGCACCTC CTACCCTCCC ACCAGAACGC ACCCCACTGG GAGAGCTGGT GGTGCAGCCT 14410 TCCCCTCCCT GTATAAGACA CTTTGCCAAG GCTCTCCCCT CTCGCCCCAT CCCTGCTTGC 14470 CCGCTCCCAC AGCTTCCTGA GGGCTAATTC TGGGAAGGGA GAGTTCTTTG CTGCCCCTGT 14530 CTGGAAGACG TGGCTCTGGG TGAGGTAGGC GGGAAAGGAT GGAGTGTTTT AGTTCTTGGG 14590 GGAGGCCACC CCAAACCCCA GCCCCAACTC CAGGGGCACC TATGAGATGG CCATGCTCAA 14650 CCCCCTCCC AGACAGGCCC TCCCTGTCTC CAGGGCCCCC ACCGAGGTTC CCAGGGCTGG 14710 14770 GCCACACCCA GGAAGGGAAA GCGGGCAGCC CCGTTTTGGG GACGTGAACG TTTTAATAAT TTTTGCTGAA TTCTTTACAA CTAAATAACA CAGATATTCT TATAAATAAA ATTGTAAAAA 14890 .



(Sh et 82 of 91)

<u>Met</u> 1	Leu	ı Th	r Pro	Pro	<u>Le</u>	<u>Leι</u>	ı Leı	ı Leı	ı Let	Pro	<u>Leu</u>	Leu	Ser	Ala	Leu
_				~											Phe
	7											4-			Asp
				4				<i></i>					Glu		
Pro	Glr	Se	Lys	Ala	Gli	Arc	Ċýs	Glņ	Pro	Asn	Glú	, His	Asn	Cys	Leu
. 00.		*			シスノロー		,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,						1 1 1	1 Towns 10 Car	.80 -Gln
					***				C. O. I	A 11 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1					
Asp	Cys	Met	Asp	<u>3</u> G1y	Ser	Asp	Glu	G1 y	Pro	His	Cys	Arg	Glu	Leu	Gln
			~~~					1117	1				7 7 0		
			,				1 / (1					126			Leu
	200					133					1 4 0		Gln		
					TOO					155			Gly		1.00
Ser	Gln	Leu	Cys	Thr 165	Asn	Thr	Asp	Gly	Ser 170	Phe	Ile	Cys	Gly		160 Val
Glu	Gly	Tyr	Leu 180	Leu	Gln	Pro	Asp	Asn 185	Arg	Ser	Cys	Lys	Ala	175 Lys	Asn
Glu	Pro	Val		Arg	Pro	Pro	Val	Leu	Leu	Ile	Ala	Asn	190 Ser	Gla	Asn
		エフン					200					205	Ile		
	210					215					220				
~ ~ ~					230					225			Ala		240
				243					250				Thr	255	Leu
			200					765	Gly				Glu 270	His	
		213					280					205	Ala		
	230					293	Phe				300	Asp	Asp		
Phe 305	Val	Cys	Asn	Arg	Asn 310	Gly	Asp	Thr	Cys	Val 315	Thr	Leu	Leu	Asp	
Glu	Leu	Tyr	Asn	Pro 325		Gly	Ile	Ala	Leu 330	Asp	Pro	Ala	Met		320 <b>Ly</b> s
Val	Phe	Phe	Thr 340		Tyr	Gly	Gln	Ile	Pro	Lys	Val	Glu	Arg	335 Cys	Asp
Met	Asp	Gly 355		Asn	Arg	Thr	Lys	345 Leu	Val	Asp	Ser		350 Ile	Val	Phe
Pro	His 370		Ile	Thr	Leu	Asp	360 Leu	Val	Ser	Arg	Leu	365 <b>Va</b> l	Tyr	Trp	Ala
Asp		Tyr	Leu	Asp	Tyr	375 Ile	Glu	Val	Val	Asp	380 Tyr	Glu	Gly	Lys	Gly
					390					395			Tyr		400
				4 V O					4 1 N				٠.	415	
			420					425					Asn. 430		
		433					440					445	Asn		
GLU	Tyr 450	Gln	Val	Val	Thr	Arg 455	Val	Asp	Lys	Gly	Gly 460	Ala	Leu	His	Ile

FIG. 14B

## (Sh et 83 of 91)

465					Gln 470					475					480
Asp	Gln	Tyr	Gly	Lys 485	Pro	Gly	Gly	Cys	Ser 490	Asp	Ile	Cys	Leu	Leu 495	Ala
Asn	Ser	His	Lys 500	Ala	Arg	Thr	Cys	Arg 505	Cys	Arg	Ser	Gly	Phe 510	Ser	Leu
Gly	Ser	<b>Asp</b> 515	Gly	Lys	Ser	Cys	Lys 520	Lys	Pro	Glu	His	Glu 525	Leu	Phe	Leu
Val	Tyr 530	Gly	Lys	Gly	Arg	Pro 535	Gly	Ile	Ile	Arg	Gly 540	Met	Asp	Met	Gly
Ala 545	Lys	Val	Pro	Asp	Glu 550	His	Met	Ile	Pro	Ile 555	Glu	Asn	Leu	Met	Asn 560
	_			565	Phe				570	_			-	575	
_			580		Leu			585		_			590		
		595			Lys		600					605	_		
Val	610				Asp	615			_		620	_	_		
Lys 625	•				Ala 630				-	635					640
				645	Lys				650	_				655	
			660	•	Met			665	-				670		
_		675			Arg		680					685			
-	690				Thr	695					700				
705					Ala 710	, ,	_		_	715		-			720
_	_			725	Ile				730					735	
-		_	740		Leu			745		_		_	750	•	
	-	755			Thr		760	_		_		765			
	77Ō	_		_	Gly	775					780				
785	_				Phe 790			_		795	_				800
		_		805	Lys				810					815	
	_		820					825		_	•		830		Asp
		835	_				840					845			Týr
													Ala	Asn	Ser
											*860				
				T. 112 J	870			Cys		875			ASP		Leu 880
Asp	Asn	Ser	ASD	Glu									Thr	Cvs	Pro
3	Y			885			第多	\$ .15	890					895	
Ser	<u>Asp</u>				` <u>Cvs</u>	<u> Glu</u>	[©] <u>Asn</u>				Tle	Pro			Trp
			.900										910		****
Leu	<u>cys</u>	915	GTA	( <u>ASD</u>	ASN	ASD	<u>278</u> 920	ענט	ASI	:- <u>ser</u>	.010	925	<u>610</u>	<u> ser</u>	Asn

FIG. 14B

## (Sheet 84 of 91)

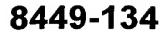
<u>Ala</u>	<u>Thr</u> 930	Cys	<u>Ser</u>	<u>Ala</u>	<u>Ara</u>	<i>Thr</i> 935	Cys	Pro	Pro	Asn	Gln	Phe	Ser	Cys: A	la
Ser	Gly	Arg	Cys	Ile	Pro		Ser	Tro	Thr	Ĉve	,940 Aen	() Jan	i	Asp A	
945					:950					955				q	60
Cys	Gly	Asp	Arg	Ser	Asp	Glu	Ser	Ala	Ser	Cys	Ala	Tyr	Pro	Thr Č	γs
		•		. 965.					970		1.72			975	12.0
File	. FTO	ren	980	<u>our</u> Oktob	Pne	<u>inr</u>	CVS	<u>ASD</u>	<u>Asn</u>	GLY	Arq	<u>Cys</u>	<u> 11e</u>	Asn T	<u>le</u>
Asn	Trp	Arq	· <u>Cys</u>	Asp	Asn	Asp	Asn	Asp	Cvs	Giv	Asp	Asn	Ser	<u>Asp</u> ;G	
<u>.</u>	. •	995			35		1000:		- 1			1005		<b>公</b> 经	
		<u>Cys</u>	<u>Ser</u>	<u>His</u>	Ser	Cys	Ser	Ser	Thr	Gln	Phe	Lys	Cys	Asn S	er
•	1010	Cic	Tla	Pro		1015			1 To		1020				
025	Arg	Cys	116		1030	nis.		SINE	Cys	( <i>ASP</i>	GIY	ASP	Asn	Aspic	<i>ys</i>
	Asp	Tyr	Ser	Asp	iGl u	Thr	His	Ala	Asn	Cvs	Thr	Asn	Gln	Ala	終
:'. · · · · · ·	•			1045				$\widetilde{\mathcal{N}}_{C}$ .	1050	•				L0557⊗	<b>*</b>
Arg	Pro	Pro	Gly	Gly.	<u>Cys</u>	<u>His</u>			<u>Glu</u>	<u>Phe</u>	<u>Gln</u>	<u>Cys</u>	Arg	Leugh	SD
; : 61 u	Ť OU	C	1060					L065	KHAN.		(grain)		1070	178	
GIY	<u>reu</u>	<u>575</u>	116	FIO	<u>Leu</u>	Arg	TED	Arg	Cys'	ASD	GIV	Asp.	Thr	Asp C	<u>Vs</u>
Met	Asp	Ser	Ser	Asp	Glu	Lvs	Ser	Cvs	Giù	Glv	٧a٦	Thr	His	Val (C	
2	1090					1095		•			1100				
Asp	Pro	Ser	Val	Lys	Phe	Gly	Cys	Lys	Asp	Ser	Ala	Arg	.Cys	Ile S	er
105	215		17- 1		1110	<b>61</b>				1115		_		11	
Lys	Ala	TP		125	Asp	G1y	Asp		Asp 1130	Cys	Glu	Asp		<i>Ser'A</i> 135	SP (*)
Glu	Gl u	Asn	Cys	Glu	Ser	Leu	Ala	Cys	<u>Arq</u>	Pro	<u>Pro</u>			Pro C	<u>vs</u>
Ala	Asn	Asn	Thr	Ser	Val	Cvs	Len	Pro	Pro	Asn	Lve		1150 Cvs	Asp G	
	]	155		•		7	160			<u> </u>	. <u> </u>	165	CYS	Vab G	<u> </u>
<u>Asn</u>	<u>Asp</u> 170	Asp	<u>Cys</u>	<u>Glv</u>	<u>Asp</u>	<u>Gly</u>	Ser	<u>Asp</u>	<u>Glu</u>		<u>Glu</u> 180	Leu	Cys	Asp G	ln
Cys	Ser	Leu	Asn	Asn	Gly	Gly	Cys	Ser	His	Asn	Cys	Ser	Val	Ala P	ro
185				3	1190				]	1195				12	00
			1	.205				]	1210				3	Gly P .215	
Asp	Asn	His	Thr 220	Cys	Gln	Ile		Ser 1225	Tyr	Cys	Ala			Leu L	ys
Cys	Ser			Cvs	Asp	Gln			Phe	Ser	Val	T.VS	L230 Cvs	Ser C	vs
	]	.235				1	L240				]	245			
Tyr	Glu	Gly	Trp	Val	Leu	Glu	Pro	Asp	Gly			Cys	Arg	Ser L	eu
	250		7	Desa		255	<b>71</b> -	Db -	0		1260				
265	PIO	Pne	Lys	PFO 1	270	TTE	116	Phe		Asn 1275	Arg	Hls	Glu	Ile A	rg 80
	Ile	Asp	Leu	His		Gly	Asp		Ser		Leu	Val		Gly L	eu
Arg	Asn	Thr		.285 Ala	Len	Asp	Phe		1290	Sar	Gla	Sor		.295 Leu T	
		)	300				. 1	305				]	1310		
Trp	Thr	Asp	Val	Val	Glu	Asp	Lys	Ile	Tyr	Arg	Gly	Lys	Leu	Leu A	sp
N c n		315	<b>T</b>	en la ca	0		1320	••-				.325			
vol!	.330	wra	rea	Thr	ser 1	2335	GIU	vai	val		Gln 1340	Tyr	Gly	Leu A	ıa
		Glu	Glv	Leu			Aso	Tro	Ile			Asn	Ile	Tyr T	rp
345				1	1350				. 3	1355			•	13	60
Val	Glu	Ser		Leu .365	Asp	Gln	Ile		Val 1370	Ala	Lys	Leu		Gly T 1375	hr
Leu	Arg		Thr		Leu	Ala		Asp		Glu	His	Pro		Ala I	le
		1	380		•		3	1385					1390		

#### (Sh t 85 of 91)

Ala Leu Asp Pro Arg Asp Gly Ile Leu Phe Trp Thr Asp Trp Asp Ala 1395 1400 Ser Leu Pro Arg Ile Glu Ala Ala Ser Met Ser Gly Ala Gly Arg Arg 1415 1420 Thr Val His Arg Glu Thr Gly Ser Gly Gly Trp Pro Asn Gly Leu Thr 1430 1435 Val Asp Tyr Leu Glu Lys Arg Ile Leu Trp Ile Asp Ala Arg Ser Asp 1450 Ala Ile Tyr Ser Ala Arg Tyr Asp Gly Ser Gly His Met Glu Val Leu 1460 1465 1470 Arg Gly His Glu Phe Leu Ser His Pro Phe Ala Val Thr Leu Tyr Gly 1480 1485 Gly Glu Val Tyr Trp Thr Asp Trp Arg Thr Asn Thr Leu Ala Lys Ala 1495 1500 Asn Lys Trp Thr Gly His Asn Val Thr Val Val Gln Arg Thr Asn Thr 1510 1515 Gln Pro Phe Asp Leu Gln Val Tyr His Pro Ser Arg Gln Pro Met Ala 1525 1530 Pro Asn Pro Cys Glu Ala Asn Gly Gly Gln Gly Pro Cys Ser His Leu 1540 1545 1550 Cys Leu Ile Asn Tyr Asn Arg Thr Val Ser Cys Ala Cys Pro His Leu 1560 Met Lys Leu His Lys Asp Asn Thr Thr Cys Tyr Glu Phe Lys Lys Phe 1570 1575 1580 Leu Leu Tyr Ala Arg Gln Met Glu Ile Arg Gly Val Asp Leu Asp Ala 1590 1595 Pro Tyr Tyr Asn Tyr Ile Ile Ser Phe Thr Val Pro Asp Ile Asp Asn 1610 Val Thr Val Leu Asp Tyr Asp Ala Arg Glu Gln Arg Val Tyr Trp Ser 1625 Asp Val Arg Thr Gln Ala Ile Lys Arg Ala Phe Ile Asn Gly Thr Gly 1635 1640 1645 Val Glu Thr Val Val Ser Ala Asp Leu Pro Asn Ala His Gly Leu Ala 1655 1660 Val Asp Trp Val Ser Arg Asn Leu Phe Trp Thr Ser Tyr Asp Thr Asn 1670 1675 1680 Lys Lys Gln Ile Asn Val Ala Arg Leu Asp Gly Ser Phe Lys Asn Ala 1685 1690 Val Val Gln Gly Leu Glu Gln Pro His Gly Leu Val Val His Pro Leu 1700 1705 1710 Arg Gly Lys Leu Tyr Trp Thr Asp Gly Asp Asn Ile Ser Met Ala Asn 1720 Met Asp Gly Ser Asn Arg Thr Leu Leu Phe Ser Gly Gln Lys Gly Pro 1735 Val Gly Leu Ala Ile Asp Phe Pro Glu Ser Lys Leu Tyr Trp Ile Ser 1750 1755 Ser Gly Asn His Thr Ile Asn Arg Cys Asn Leu Asp Gly Ser Gly Leu 1765 1770 Glu Val Ile Asp Ala Met Arg Ser Gln Leu Gly Lys Ala Thr Ala Leu 1785 1790 Ala Ile Met Gly Asp Lys Leu Trp Trp Ala Asp Gln Val Ser Glu Lys 1800 . Met Gly Thr Cys Ser Lys Ala Asp Gly Ser Gly Ser Val Val Leu Arg 1810 1815 . 1820 Asn Ser Thr Thr Leu Val Met His Met Lys Val Tyr Asp Glu Ser Ile 1830 1835 Gln Leu Asp His Lys Gly Thr Asn Pro Cys Ser Val Asn Asn Gly Asp 1845 1850 Cys Ser Gln Leu Cys Leu Pro Thr Ser Glu Thr Thr Arg Ser Cys Met

#### (Sheet 86 of 91)

Cys Thr Ala Gly Tyr Ser Leu Arg Ser Gly Gln Gln Ala Cys Glu Gly 1865 1880 Val Gly Ser Phe Leu Leu Tyr Ser Val His Glu Gly Ile Arg Gly Ile 1895 Pro Leu Asp Pro Asn Asp Lys Ser Asp Ala Leu Val Pro Val Ser Gly Thr Ser Leu Ala Val Gly Ile Asp Phe His Ala Glu Asn Asp Thr Ile Tyr Trp Val Asp Met Gly Leu Ser Thr Ile Ser Arg Ala Lys Arg Asp 1930 1945 Gln Thr Trp Arg Glu Asp Val Val Thr Asn Gly Ile Gly Arg Val Glu 1960 Gly Ile Ala Val Asp Trp Ile Ala Gly Asn Ile Tyr Trp Thr Asp Gln 1975 Gly Phe Asp Val Ile Glu Val Ala Arg Leu Asn Gly Ser Phe Arg Tyr Val Val Ile Ser Gln Gly Leu Asp Lys Pro Arg Ala Ile Thr Val His 2010 Pro Glu Lys Gly Tyr Leu Phe Trp Thr Glu Trp Gly Gln Tyr Pro Arg 2025 Ile Glu Arg Ser Arg Leu Asp Gly Thr Glu Arg Val Val Leu Val Asn 2040 Val Ser Ile Ser Trp Pro Asn Gly Ile Ser Val Asp Tyr Gln Asp Gly 2055 Lys Leu Tyr Trp Cys Asp Ala Arg Thr Asp Lys Ile Glu Arg Ile Asp 2060 2075 Leu Glu Thr Gly Glu Asn Arg Glu Val Val Leu Ser Ser Asn Asn Met 2090 Asp Met Phe Ser Val Ser Val Phe Glu Asp Phe Ile Tyr Trp Ser Asp 2105 Arg Thr His Ala Asn Gly Ser Ile Lys Arg Gly Ser Lys Asp Asn Ala 2120 Thr Asp Ser Val Pro Leu Arg Thr Gly Ile Gly Val Gln Leu Lys Asp 2135 Ile Lys Val Phe Asn Arg Asp Arg Gln Lys Gly Thr Asn Val Cys Ala 2140 2150 2155 Val Ala Asn Gly Gly Cys Gln Gln Leu Cys Leu Tyr Arg Gly Arg Gly 2170 Gln Arg Ala Cys Ala His Gly Met Leu Ala Glu Asp Gly Ala 2185 Ser Cys Arg Glu Tyr Ala Gly Tyr Leu Leu Tyr Ser Glu Arg Thr Ile 2200 Leu Lys Ser Ile His Leu Ser Asp Glu Arg Asn Leu Asn Ala Pro Val 2215 Gln Pro Phe Glu Asp Pro Glu His Met Lys Asn Val Ile Ala Leu Ala 2220 2230 · 2235 Phe Asp Tyr Arg Ala Gly Thr Ser Pro Gly Thr Pro Asn Arg Ile Phe 2250 Phe Ser Asp Ile His Phe Gly Asn Ile Gln Gln Ile Asn Asp Asp Gly 2265 Ser Arg Arg Ile Thr Ile Val Glu Asn Val Gly Ser Val Glu Gly Leu 2280 2285 Ala Tyr His Arg Gly Trp Asp Thr Leu Tyr Trp Thr Ser Tyr Thr Thr 2295 2300 Ser Thr Ile Thr Arg His Thr Val Asp Gln Thr Arg Pro Gly Ala Phe 2310 Glu Arg Glu Thr Val Ile Thr Met Ser Gly Asp Asp His Pro Arg Ala 2315 2325 2330



#### (Sh et 87 of 91)

Phe Val Leu Asp Glu Cys Gln Asn Leu Met Phe Trp Thr Asn Trp Asn Glu Gln His Pro Ser Ile Met Arg Ala Ala Leu Ser Gly Ala Asn Val 2345 2360 Leu Thr Leu Ile Glu Lys Asp Ile Arg Thr Pro Asn Gly Leu Ala Ile Asp His Arg Ala Glu Lys Leu Tyr Phe Ser Asp Ala Thr Leu Asp Lys Ile Glu Arg Cys Glu Tyr Asp Gly Ser His Arg Tyr Val Ile Leu Lys Ser Glu Pro Val His Pro Phe Gly Leu Ala Val Tyr Gly Glu His Ile 2425 Phe Trp Thr Asp Trp Val Arg Arg Ala Val Gln Arg Ala Asn Lys His 2440 Val Gly Ser Asn Met Lys Leu Leu Arg Val Asp Ile Pro Gln Gln Pro 2455 Met Gly Ile Ile Ala Val Ala Asn Asp Thr Asn Ser Cys Glu Leu Ser Pro Cys Arg Ile Asn Asn Gly Gly Cys Gln Asp Leu Cys Leu Leu Thr His Gln Gly His Val Asn Cys Ser Cys Arg Gly Gly Arg Ile Leu Gln 2490 2505 Asp Asp Leu Thr Cys Arg Ala Val Asn Ser Ser Cys Arg Ala Gln Asp 2520 Glu Phe Glu Cys Ala Asn Gly Glu Cys Ile Asn Phe Ser Leu Thr Cys Asp Gly Val Pro His Cys Lys Asp Lys Ser Asp Glu Lys Pro Ser Tyr 2540 Cys Asn Ser Arg Arg Cys Lys Lys Thr Phe Arg Gln Cys Ser Asn Gly 2555 2570 Arg Cys Val Ser Asn Met Leu Trp Cys Asn Gly Ala Asp Asp Cys Gly 2585 Asp Gly Ser Asp Glu Ile Pro Cys Asn Lys Thr Ala Cys Gly Val Gly .2600 Glu Phe Arg Cys Arg Asp Gly Thr Cys Ile Gly Asn Ser Ser Arg Cys 2615 Asn Gln Phe Val Asp Cys Glu Asp Ala Ser Asp Glu Met Asn Cys Ser 2635 Ala Thr Asp Cys Ser Ser Tyr Phe Arg Leu Gly Val Lys Gly Val Leu Phe Gln Pro Cys Glu Arg Thr Ser Leu Cys Tyr Ala Pro Ser Trp Val 2650 2665 Cys Asp Gly Ala Asn Asp Cys Gly Asp Tyr Ser Asp Glu Arg Asp Cys 2680 Pro Gly Val Lys Arg Pro Arg Cys Pro Leu Asn Tyr Phe Ala Cys Pro 2695 Ser Gly Arg Cys Ile Pro Met Ser Trp Thr Cys Asp Lys Glu Asp Asp 2710 2715 Cys Glu His Gly Glu Asp Glu Thr His Cys Asn Lys Phe Cys Ser Glu 2730 Ala Gln Phe Glu Cys Gln Asn His Arg Cys Ile Ser Lys Gln Trp Leu 2745 Cys Asp Gly Ser Asp Asp Cys Gly Asp Gly Ser Asp Glu Ala Ala His 2760 Cys Glu Gly Lys Thr Cys Gly Pro Ser Ser Phe Ser Cys Pro Gly Thr 2765 His Val Cys Val Pro Glu Arg Trp Leu Cys Asp Gly Asp Lys Asp Cys 2780 Ala Asp Gly Ala Asp Glu Ser Ile Ala Ala Gly Cys Leu Tyr Asn Ser 2795

#### (Sh t 88 of 91)

2805 2810 Thr Cys Asp Asp Arg Glu Phe Met Cys Gln Asn Arg Gln Cys Ile Pro 2825 Lys His Phe Val Cys Asp His Asp Arg Asp Cys Ala Asp Gly Ser Asp 2840 Glu Ser Pro Glu Cys Glu Tyr Pro Thr Cys Gly Pro Ser Glu Phe Arg 2855 2860 Cys Ala Asn Gly Arg Cys Leu Ser Ser Arg Gln Trp Glu Cys Asp Gly 865 2870 2875 Glu Asn Asp Cys His Asp Gln Ser Asp Glu Ala Pro Lys Asn Pro His 2885 2890 Cys Thr Ser Pro Glu His Lys Cys Asn Ala Ser Ser Gln Phe Leu Cys 2900 2905 Ser Ser Gly Arg Cys Val Ala Glu Ala Leu Leu Cys Asn Gly Gln Asp 2920 2925 Asp Cys Gly Asp Ser Ser Asp Glu Arg Gly Cys His Ile Asn Glu Cys 2935 Leu Ser Arg Lys Leu Ser Gly Cys Ser Gln Asp Cys Glu Asp Leu Lys 2950 2955 Ile Gly Phe Lys Cys Arg Cys Arg Pro Gly Phe Arg Leu Lys Asp Asp 2965 2970 Gly Arg Thr Cys Ala Asp Val Asp Glu Cys Ser Thr Thr Phe Pro Cys 2985 Ser Gln Arg Cys Ile Asn Thr His Gly Ser Tyr Lys Cys Leu Cys Val 3000 3005 Glu Gly Tyr Ala Pro Arg Gly Gly Asp Pro His Ser Cys Lys Ala Val 3015 3020 Thr Asp Glu Glu Pro Phe Leu Ile Phe Ala Asn Arg Tyr Tyr Leu Arg 3030 3035 Lys Leu Asn Leu Asp Gly Ser Asn Tyr Thr Leu Leu Lys Gln Gly Leu 3045 3050 Asn Asn Ala Val Ala Leu Asp Phe Asp Tyr Arg Glu Gln Met Ile Tyr 3060 3065 Trp Thr Asp Val Thr Thr Gln Gly Ser Met Ile Arg Arg Met His Leu 3080 3085 Asn Gly Ser Asn Val Gln Val Leu His Arg Thr Gly Leu Ser Asn Pro 3095 3100 Asp Gly Leu Ala Val Asp Trp Val Gly Gly Asn Leu Tyr Trp Cys Asp 3110 3115 3120 Lys Gly Arg Asp Thr Ile Glu Val Ser Lys Leu Asn Gly Ala Tyr Arg 3125 3130 Thr Val Leu Val Ser Ser Gly Leu Arg Glu Pro Arg Ala Leu Val Val 3140 3145 Asp Val Gln Asn Gly Tyr Leu Tyr Trp Thr Asp Trp Gly Asp His Ser 3160 3165 Leu Ile Gly Arg Ile Gly Met Asp Gly Ser Ser Arg Ser Val Ile Val 3175 3180 Asp Thr Lys Ile Thr Trp Pro Asn Gly Leu Thr Leu Asp Tyr Val Thr 3190 3195 Glu Arg Ile Tyr Trp Ala Asp Ala Arg Glu Asp Tyr Ile Glu Phe Ala 3205 3210 Ser Leu Asp Gly Ser Asn Arg His Val Val Leu Ser Gln Asp Ile Pro 3220 3225 His Ile Phe Ala Leu Thr Leu Phe Glu Asp Tyr Val Tyr Trp Thr Asp 3240 3245 Trp Glu Thr Lys Ser Ile Asn Arg Ala His Lys Thr Thr Gly Thr Asn 3255 Lys Thr Leu Leu Ile Ser Thr Leu His Arg Pro Met Asp Leu His Val 3270 3275

### 8449-134 (Sh et 89 of 91)

Phe His Ala Leu Ary Gin Pro Asp Val Pro Asn His Pro Cys Lys Val 3285 3290 Asn Asn Gly Gly Cys Ser Asn Leu Cys Leu Leu Ser Pro Gly Gly Gly 3300 3305 His Lys Cys Ala Cys Pro Thr Asn Phe Tyr Leu Gly Ser Asp Gly Arg 3320 3325 Thr Cys Val Ser Asn Cys Thr Ala Ser Gln Phe Val Cys Lys Asn Asp 3335 3340 Lys Cys Ile Pro Phe Trp Trp Lys Cys Asp Thr Glu Asp Asp Cys Gly 3350 3355 Asp His Ser Asp Glu Pro Pro Asp Cys Pro Glu Phe Lys Cys Arg Pro 3365 3370 Gly Gln Phe Gln Cys Ser Thr Gly Ile Cys Thr Asn Pro Ala Phe Ile 3380 3385 Cys Asp Gly Asp Asn Asp Cys Gln Asp Asn Ser Asp Glu Ala Asn Cys. 3400 Asp Ile His Val Cys Leu Pro Ser Gln Phe Lys Cys Thr Asn Thr Asn 3405 3415 3420 Arg Cys Ile Pro Gly Ile Phe Arg Cys Asn Gly Gln Asp Asn Cys Gly 3430 3435 Asp Gly Glu Asp Glu Arg Asp Cys Pro Glu Val Thr Cys Ala Pro Asn 3445 3450 3455 Gln Phe Gln Cys Ser Ile Thr Lys Arg Cys Ile Pro Arg Val Trp Val 3460 3465 Cys Asp Arg Asp Asn Asp Cys Val Asp Gly Ser Asp Glu Pro Ala Asn 3480 3485 Cys Thr Gln Met Thr Cys Gly Val Asp Glu Phe Arg Cys Lys Asp Ser 3495 3500 Gly Arg Cys Ile Pro Ala Arg Trp Lys Cys Asp Gly Glu Asp Asp Cys 3510 3515 Gly Asp Gly Ser Asp Glu Pro Lys Glu Glu Cys Asp Glu Arg Thr Cys 3525 3530 Glu Pro Tyr Gln Phe Arg Cys Lys Asn Asn Arg Cys Val Pro Gly Arg 3545 Trp Gln Cys Asp Tyr Asp Asn Asp Cys Gly Asp Asn Ser Asp Glu Glu 3555 3565 Ser Cys Thr Pro Arg Pro Cys Ser Glu Ser Glu Phe Ser Cys Ala Asn 3575 3580 Gly Arg Cys Ile Ala Gly Arg Trp Lys Cys Asp Gly Asp His Asp Cys 3590 3595 Ala Asp Gly Ser Asp Glu Lys Asp Cys Thr Pro Arg Cys Asp Met Asp 3610 Gln Phe Gln Cys Lys Ser Gly His Cys Ile Pro Leu Arg Trp Arg Cys 3620 3625 3630 Asp Ala Asp Ala Asp Cys Met Asp Gly Ser Asp Glu Glu Ala Cys Gly 3635 3640 3645 Thr Gly Val Arg Thr Cys Pro Leu Asp Glu Phe Gln Cys Asn Asn Thr 3655 . 3660 Leu Cys Lys Pro Leu Ala Trp Lys Cys Asp Gly Glu Asp Asp Cys Gly 3670 3675 Asp Asn Ser Asp Glu Asn Pro Glu Glu Cys Ala Arg Phe Val Cys Pro 3685 3690 3695 Pro Asn Arg Pro Phe Arg Cys Lys Asn Asp Arg Val Cys Leu Trp Ile 3700 3705 3710 Gly Arg Gln Cys Asp Gly Thr Asp Asn Cys Gly Asp Gly Thr Asp Glu 3715 3720 3725 Glu Asp Cys Glu Pro Pro Thr Ala His Thr Thr His Cys Lys Asp Lys Lys Glu Phe Leu Cys Arg Asn Gln Arg Cys Leu Ser Ser Leu Arg

#### (Sh et 90 of 91)

3750 Cys Asn Met Phe Asp Asp Cys Gly Asp Gly Ser Asp Glu Glu Asp Cys 3755 3765 3770 Ser Ile Asp Pro Lys Leu Thr Ser Cys Ala Thr Asn Ala Ser Ile Cys 3780 3785 Gly Asp Glu Ala Arg Cys Val Arg Thr Glu Lys Ala Ala Tyr Cys Ala 3800 Cys Arg Ser Gly Phe His Thr Val Pro Gly Gln Pro Gly Cys Gln Asp 3815 3820 Ile Asn Glu Cys Leu Arg Phe Gly Thr Cys Ser Gln Leu Cys Asn Asn 3830 3835 Thr Lys Gly Gly His Leu Cys Ser Cys Ala Arg Asn Phe Met Lys Thr 3845 3850 His Asn Thr Cys Lys Ala Glu Gly Ser Glu Tyr Gln Val Leu Tyr Ile 3865 Ala Asp Asp Asn Glu Ile Arg Ser Leu Phe Pro Gly His Pro His Ser 3875 3880 Ala Tyr Glu Gln Ala Phe Gln Gly Asp Glu Ser Val Arg Ile Asp Ala 3895 3900 Met Asp Val His Val Lys Ala Gly Arg Val Tyr Trp Thr Asn Trp His 3910 3915 Thr Gly Thr Ile Ser Tyr Arg Ser Leu Pro Pro Ala Ala Pro Pro Thr 3925 3930 Thr Ser Asn Arg His Arg Arg Gln Ile Asp Arg Gly Val Thr His Leu 3945 3950 Asn Ile Ser Gly Leu Lys Met Pro Arg Gly Ile Ala Ile Asp Trp Val 3960 3965 Ala Gly Asn Val Tyr Trp Thr Asp Ser Gly Arg Asp Val Ile Glu Val 3975 3980 Ala Gln Met Lys Gly Glu Asn Arg Lys Thr Leu Ile Ser Gly Met Ile 3990 3995 Asp Glu Pro His Ala Ile Val Val Asp Pro Leu Arg Gly Thr Met Tyr 4005 4010 Trp Ser Asp Trp Gly Asn His Pro Lys Ile Glu Thr Ala Ala Met Asp 4025 4030 Gly Thr Leu Arg Glu Thr Leu Val Gln Asp Asn Ile Gln Trp Pro Thr 4035 4040 Gly Leu Ala Val Asp Tyr His Asn Glu Arg Leu Tyr Trp Ala Asp Ala 4055 4060 Lys Leu Ser Val Ile Gly Ser Ile Arg Leu Asn Gly Thr Asp Pro Ile 4070 4075 Val Ala Ala Asp Ser Lys Arg Gly Leu Ser His Pro Phe Ser Ile Asp 4085 4090 Val Phe Glu Asp Tyr Ile Tyr Gly Val Thr Tyr Ile Asn Asn Arg Val Phe Lys Ile His Lys Phe Gly His Ser Pro Leu Val Asn Leu Thr Gly 4115 4120 4125 Gly Leu Ser His Ala Ser Asp Val Val Leu Tyr His Gln His Lys Gln 4135 4140 Pro Glu Val Thr Asn Pro Cys Asp Arg Lys Lys Cys Glu Trp Leu Cys 4150 4155 Leu Leu Ser Pro Ser Gly Pro Val Cys Thr Cys Pro Asn Gly Lys Arg 4165 4170 Leu Asp Asn Gly Thr Cys Val Pro Val Pro Ser Pro Thr Pro Pro Pro 4180 4185 4190 Asp Ala Pro Arg Pro Gly Thr Cys Asn Leu Gln Cys Phe Asn Gly Gly 4200 Ser Cys Phe Leu Asn Ala Arg Arg Gln Pro Lys Cys Arg Cys Gln Pro

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#### 8449-134

#### (She t 91 of 91)

Arg Tyr Thr Gly Asp Lys Cys Glu Leu Asp Gln Cys Trp Glu His Cys 4230 Arg Asn Gly Gly Thr Cys Ala Ala Ser Pro Ser Gly Met Pro Thr Cys 4245 4250 Arg Cys Pro Thr Gly Phe Thr Gly Pro Lys Cys Thr Gln Gln Val Cys 4265 Ala Gly Tyr Cys Ala Asn Asn Ser Thr Cys Thr Val Asn Gln Gly Asn 4275 4280 4285 Gln Pro Gln Cys Arg Cys Leu Pro Gly Phe Leu Gly Asp Arg Cys Gln 4295 4300 Tyr Arg Gln Cys Ser Gly Tyr Cys Glu Asn Phe Gly Thr Cys Gln Met 305 4310 4315 Ala Ala Asp Gly Ser Arg Gln Cys Arg Cys Thr Ala Tyr Phe Glu Gly 4325 4330 Ser Arg Cys Glu Val Asn Lys Cys Ser Arg Cys Leu Glu Gly Ala Cys 4345 4350 Val Val Asn Lys Gln Ser Gly Asp Val Thr Cys Asn Cys Thr Asp Gly 4360 4365 Arg Val Ala Pro Ser Cys Leu Thr Cys Val Gly His Cys Ser Asn Gly 4375 Gly Ser Cys Thr Met Asn Ser Lys Met Met Pro Glu Cys Gln Cys Pro 4390 4395 Pro His Met Thr Gly Pro Arg Cys Glu Glu His Val Phe Ser Gln Gln 4405 4410 4415 Gln Pro Gly His Ile Ala Ser Ile Leu Ile Pro Leu Leu Leu Leu 4425 4430 Leu Leu Val Leu Val Ala Gly Val Val Phe Trp Tyr Lys Arg Arg Val 4435 4440 Gln Gly Ala Lys Gly Phe Gln His Gln Arg Met Thr Asn Gly Ala Met 4455 4460 Asn Val Glu Ile Gly Asn Pro Thr Tyr Lys Met Tyr Glu Gly Gly Glu 4470 4475 Pro Asp Asp Val Gly Gly Leu Leu Asp Ala Asp Phe Ala Leu Asp Pro 4485 4490 4495 Asp Lys Pro Thr Asn Phe Thr Asn Pro Val Tyr Ala Thr Leu Tyr Met 4500 4505 Gly Gly His Gly Ser Arg His Ser Leu Ala Ser Thr Asp Glu Lys Arg 4520 4525 Glu Leu Leu Gly Arg Gly Pro Glu Asp Glu Ile Gly Asp Pro Leu Ala 4535